We claim:

1. A histone deacetylase inhibitor of formula (1):

or a pharmaceutically acceptable salt thereof, wherein

Ar² is a saturated or mono- or poly- unsaturated C₅-Ci₄-mono- or fused poly- cyclic hydrocarbyl, optionally containing one, two, three, or four annular heteroatoms per ring optionally substituted with one or more groups selected from C_r C_T-alkyl, hydroxy, C_r C_T-alkoxy, halo, and amino, provided that an annular O or S is not adjacent to another annular O or S;

R⁵ and R⁶ are independently selected from the group consisting of hydrogen, C_r C₇ alkyl, aryl, and aralkyl;

R², R³ and R⁴ are independently selected from the group consisting of hydrogen, halogen, -NH₂, nitro, hydroxy, aryl, heterocyclyl, C₃-C₈-cycloalkyl, heteroaryl, C_r C₇-akyl, haloalkyl, C_r C₇-alkenyl, C_T-C₇ alkynyl, C_r C₇-acyl, CrC ₇-alkyl-aryloxy, Ci-C₇-alkyl-arylsulfanyl, Ci-C₇-alkyl-arylsulfinyl, C_r C₇-alkyl-arylaminosulfonyl, Ci-C₇-alkyl-arylamine, CrC ₇-alkynyl-C(O)-amine, C₁-C₇ alkenyl-C(O)-amine, Ci-C₇-alkynyl-R⁹, C_r C₇-alkenyl-R⁹ wherein R⁹ is hydrogen , hydroxy, amino, C₁-Cralkyl or C_r C₇-alkoxy;

q is O or 1;

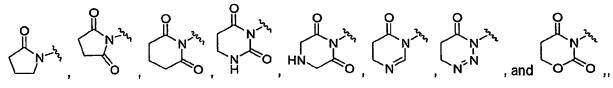
R¹ is a mono-, bi-, or tri-cyclic aryl or heteroaryl, each of which is optionally substituted; Y is any pharmaceutically acceptable chemical moiety consisting of 1 to 50 atoms; and provided that

when R^1 is /V-imidazolyl, R^2 - R^4 are H, q is 0, and Ar^2 is pyridine, Y is not CI; and when R^1 is p-aminophenyl, R^2 - R^4 are H, q is 0, and Ar^2 is phenyl, Y is not H.

- 2. The compound according to claim 1 wherein R1 is phenyl, naphthyl, anthracenyl, or fluorenyl.
- 3. The compound according to claim 1 wherein R¹ is furanyl or thienyl.
- 4. The compound according to claim 2 wherein R2, R3, and R4 are all -H.

5. The compound according to claim 3 wherein R2, R3, and R4 are all -H.

- 6. The compound according to claim 1 wherein Y is Cy2-X1- and
- Cy² is hydrogen, cycloalkyl, aryl, heteroaryl, or heterocyclyl, each of which is optionally substituted and each of which is optionally fused to one or two aryl or heteroaryl rings, or to one or two saturated or partially unsaturated cycloalkyl or heterocyclic rings, and wherein any of the aforementioned rings are optionally substituted; and
- X¹ is selected from the group consisting of a covalent bond, M¹-IA W¹, and L²-M²-L² wherein
- L^2 , at each occurrence, is independently selected from the group consisting of a chemical bond, C_0 - C_4 -hydrocarbyl, C_0 - C_4 -hydrocarbyl-(NH)- C_0 - C_4 -hydrocarbyl, C_0 - C_4 -hydrocarbyl, and C_0 - C_4 -hydrocarbyl-(0)- C_0 - C_4 -hydrocarbyl, provided that L^2 is not a chemical bond when X^1 is MMAM X^1 ;
- M^1 , at each occurrence, is independently selected from the group consisting of -0-, -N(R⁷)-, -S-, -S(O)-, $S(O)_2$ -, -S(O)₂N(R⁷)-, -N(R⁷J-S(O)₂-, -C(O)-NH-, -NH-C(O)-, -NH-C(O)-O-and -0-C(O)-NH-, -NH-C(O)-NH-, -NH-C(O)-NH-,
- R⁷ is selected from the group consisting of hydrogen, C_r C₆-hydrocarbyl, aryl, aralkyl, acyl, C₀-C₆-hydrocarbyl-heterocyclyl, and Co-Ce-hydrocarbyl-heteroaryl, wherein the hydrocarbyl moieties are optionally substituted with -OH, -NH₂, -N(H)CH₃, -N(CH₃)₂, or halo; and
- M² is selected from the group consisting of M¹, heteroarylene, and heterocyclylene, either of which rings optionally is substituted.
- 7. The compound according to claim 6, wherein X¹ is selected from the group consisting of a N(Z)-Co-C₇-alkyl-, -O-Co-Cy-alkyl-, -C(H)=CH-C₀-C₇-alkyl-, -S-C₀-C₇-alkyl-, or -C₇-C₇-alkyl-, wherein Z is H or -CrC₇-alkyl- optionally substituted with -OH, -NH₂, or halo.
- 8. The compound according to claim 6, wherein X¹ is selected from methylene, aminomethyl, and thiomethyl.
- 9. The compound according to claim 6, wherein Cy2 is selected from



each of which optionally is substituted and optionally is fused to one or more aryl rings.

 The compound according to claim 6 wherein Cy² is aryl or heteroaryi, each optionally substituted.

- 11. The compound according to claim 6 wherein Cy² is phenyl, pyrimidinyl, benzoimidazolyl or benzothiazolyl, each of which is optionally substituted.
- 12. The compound according to claim 11 wherein Cy² has from one and three substituents independently selected from the group consisting of Ci-Cralkoxy, halo, di-C₁-C₇-alkylamino-C₁-C₇ alkoxy and heteroaryi.
- 13. The compound according to claim 12 wherein the substituents are selected from methoxy, fluoro, chloro, pyridinyl and dimethylamino-ethoxy.
- 14. The compound according to claim 13 wherein Cy^2 is phenyl substituted with one to three $\mathrm{CH}_3\mathrm{O}$ -.
- 15. The compound according to claim 6 wherein Y is (V-L4VV-L3-, and
- L^3 is a direct bond, -d-Ce-hydrocarbyl, -(C₁-C₃-hydrocarbyl) $_{m1}$ -X'-(C₁-C₃- hydrocarbyl) $_{m2}$, -NH-(C₀-C₃- hydrocarbyl), (Ci-C₃- hydrocarbyO-NH-, or -NH-(Ci-C₃- hydrocarbyl)-NH-;

m1 and m2 are independently Oor 1;

X' is $-N(R^{21})$ -, $-C(O)N(R^{21})$ -, $N(R^{21}JC(O)$ -, -0-, or -S-;

 R^{21} is -H, V"-(Ci-C₆-hydrocarbyl) _a;

L⁴ is (Ci-C₆-hydrocarbyl) _a-IVI-(Ci-C6-hydrocarbyl)b;

a and b are independently O or 1;

M is -NH-, -NHC(O)-, -C(O)NH-, -C(O)-, -SO₂-, -NHSO₂-, or -SO₂NH-

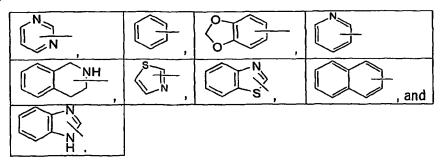
V, V, and V" are independently selected from cycloalkyl, heterocyclyl, aryl, and heteroaryi; t is O or 1.

16. The compound according to claim 15 wherein Y is V-L3 and

L³ is -NH-CH- or -CH-NH-;

V is phenyl optionally substituted with from 1 to 3 moieties independently selected from halo, hydroxy, Ci-C₆-hydrocarbyl, CrC₆-hydrocarbyl-oxy or -thio (particularly methoxy or methylthio), wherein each of the hydrocarbyl moieties are optionally substituted with one or more moieties independently selected from halo, nitroso, amino, sulfonamido, and cyano.

17. The compound according to claim 16 wherein V is an optionally substituted ring moiety selected from:



- 18. The compound according to claim 6 wherein
- Cy² is cycloalkyl, aryl, heteroaryl, or heterocyclyl, each of which optionally is substituted, and each of which optionally is fused to one or more aryl or heteroaryl rings, or to one or more saturated or partially unsaturated cycloalkyl or heterocyclic rings, each of which rings optionally is substituted, provided that when Cy² is a cyclic moiety having -C(O)-, -C(S)-, -S(O)-, or -S(O)₂- in the ring, then Cy² is not additionally substituted with a group comprising an aryl or heteroaryl ring; and
- X¹ is selected from the group consisting of a chemical bond, L³, W¹L³, IΛW¹, W¹-LΛW¹, and LAW¹-!.³, wherein
- W¹, at each occurrence, is S, O, or N(R⁹), where R⁹ is selected from the group consisting of hydrogen, alkyl, aryl, and aralkyl; and
- ${\sf L^3}$ is ${\sf C_1C_4}$ alkylene, ${\sf C_2C_4}$ alkenylene, or ${\sf C_2C_4}$ alkynylene.
- 19. The compound according to claim 6 wherein Y is selected from:
 - Ai-LrBr, wherein Ai is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein Li is -(CH₂)o-iNH(CH₂)o-r, -NHC(O)-, or -NHCH₂; and wherein Bi is phenyl or a covalent bond;
 - b) A₂-L₂-B₂-, wherein A₂ is CH₃(C=CH₂)-, optionally substituted cycloalkyl, optionally substituted aryl; wherein L₂ is -CsC-; and wherein B₂ is a covalent bond;
 - A₃-L₃-B₃-, wherein A₃ is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein L₃ is a covalent bond; and wherein B₃ is CH₂NH-;

d) A_4 - L_4 - B_4 -, wherein A_4 is an optionally substituted aryl; wherein L_4 is -NHCH $_2$ -; and wherein B_4 is a thienyl group;

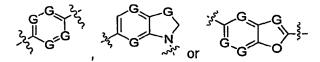
- e) $A_5-L_5-B_5$, wherein A_5 is an optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein L_5 is a covalent bond; and wherein B_5 is -SCH₂-;
- f) morpholinyl-CH 2-
- g) optionally substituted aryl;
- h) $A_6-L_6-B_6$, wherein A_6 is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein L_6 is a covalent bond; and wherein B_6 is NHCH $_2$ -;
- i) A_7 -LrB $_{\mathcal{T}}$, wherein A_7 is an optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein L_7 is a covalent bond; and wherein B_7 is -CH $_2$ -;
- j) optionally substituted heteroaryl or optionally substituted heterocyclyl;
- k) A_8 - L_8 - B_8 -, wherein A_8 is optionally substituted phenyl; wherein L_8 is a covalent bond; and wherein B_8 is -0-;
- 1) A9-L9-B9-, wherein A_9 is an optionally substituted aryl; wherein L_9 is a covalent bond; and wherein B_9 is a furan group;
- m) AiO-L₁O-BiO-, wherein Ai₀ is an optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein Li₀ is -CH(CH₂CH₃)-; and wherein Bi₀ is -NHCH₂-;
- n) $A_{11}^-L_{11}^-Bu^-$, wherein A_{11} is an optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein L_{11} is a covalent bond; and wherein B_{11} is -OCH 2^- ;
- 0) A₁₂-L₁₂-Bi₂-, wherein A₁₂ is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein L₁₂ is-NHC(O)-; and wherein B₁₂ is -N(optionally substituted aryl)CH₂-;
- p) A₁₃-L₁₃-B₁₃-, wherein A₁₃ is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein L₁₃ is a covalent bond; and wherein B₁₃ is -NHC(O)-;
- q) Ai₄-Lu-B₁₄-, wherein A₁₄ is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein L₁₄ is-NHC(0)(optionally substituted heteroaryl); and wherein B₁₄ is -S-S-;
- r) $F_3CC(O)NH-;$

s) Ai ₅I_i₅B₁₅, wherein A₁₅ is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein U s is-(CH₂)o-iNH(optionally substituted heteroaryl)-; and wherein B₁₅ is -NHCH₂-;

- t) A₁₆-L₁₆-B₁₆-, wherein Ai₆ is an optionally substituted aryl, optionally substituted heterocyclyl; wherein Li₆ is a covalent bond; and wherein Bi₆ is -N(optionally substituted alkyl)CH₂-; and
- u) A₁₇-L₁₇-B₁₇-, wherein A₁₇ is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein L₁₇ is a covalent bond; and wherein Bi₇ is -(optionally substituted aryl-CH₂)₂-N-.
- 20. The compound according to claim 6 wherein Y is selected from:
 - a) D₁-E₁-F₁, wherein Di is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein Ei is -CH₂- or a covalent bond; and wherein Fi is a covalent bond;
 - b) D_2 - E_2 - F_2 -, wherein D_2 is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein E_2 is -NH(CH $_2$) $_0$ - $_2$ -; and wherein F_2 is a covalent bond;
 - c) D3-E3-F3-, wherein D₃ is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein E₃ is -(CH₂)_ENH-; and wherein F₃ is a covalent bond;
 - d) D_4 - E_4 - F_4 -, wherein D_4 is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein E_4 is -S(CH $_2$) $_0$ - $_2$ -; and wherein F_4 is a covalent bond;
 - e) $D_5 = E_5 = F_5$, wherein D_5 is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein E_5 is -(CH $_2$) $_{0-2}$ S-; and wherein F_5 is a covalent bond; and
 - f) D_6 - E_6 - F_5 , wherein D_6 is an optionally substituted aryl, optionally substituted heteroaryl or optionally substituted heterocyclyl; wherein E_6 is -NH(CH $_2$)o- $_2$ NH-; and wherein F_6 is a covalent bond.

21. The compound according to claim 2 wherein R² to R⁴ are independently hydrogen, -NH₂, nitro, furanyl, chloro, fluoro, butyl, trifluoromethyl, bromo, thienyl, phenyl, -CHCHC(O)-NH₂, -C=CCH_TR⁹ wherein R⁹ is hydrogen, C_r C_Talkyl, hydroxy, amino, or d-C _Talkoxy.

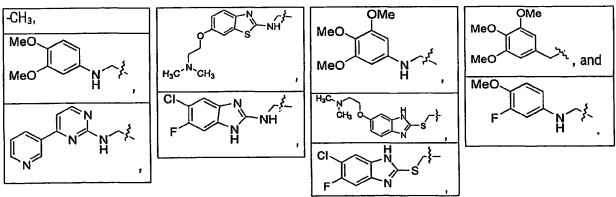
- 22. The compound according to claim 3 wherein R² to R⁴ are independently hydrogen, -NH₂, nitro, furanyl, chloro, fluoro, butyl, trifluoromethyl, bromo, thienyl, phenyl, -CHCHC(O)-NH₂, -C≡CCH₂R³ wherein R³ is hydrogen, CrC ralkyl, hydroxy, amino, or Ci-Cy-alkoxy.
- 23. The compound according to claim 6 wherein q is O and X^1 is independently selected from the group consisting of a -NH-CH₂, -S-CH₂- and -CH₂.
- 24. The compound according to claim 1 wherein Ar² has the formula



and wherein G, at each occurrence, is independently N or C, and C is optionally substituted.

25. The compound according to claim 24 wherein Ar² has the formula

- 26. The compound according to claim 24 wherein Ar² is selected from the group consisting of phenylene, benzofuranylene and indolinylene.
- 27. The compound according to claim 6 wherein the moiety formed by Cy²-x¹ is selected from:



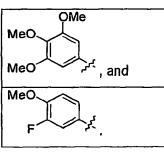
28. The compound of claim 6 of formula (2):

or a pharmaceutically acceptable salt thereof, wherein

R² and R³ are independently selected from the group consisting of hydrogen, trifluoromethyl, butyl, - (CH₂)S-OH, chloro, fluoro, amino, phenyl, thienyl, furanyl, -CHCCHC(O)NH₂, -CsCCH₂-OH, - C=CCH₂-OCH₃; and

the A ring is optionally further substituted with from 1 to 3 substituents independently selected from methyl, hydroxy, methoxy, halo, and amino.

29. The compound according to claim 28 wherein Cy2 is selected from:



- 30. The compound according to claim 28 wherein the A ring is not further substituted.
- 31. The compound according to claim 28 wherein R² and R³ are -H.
- 32. A compound according to claim 1 selected from:

 $\label{eq:local_problem} $$ $V-[2-amino-5-(2-thienyl)phenyl]-4-{[(3,4-dimethoxyphenyl)amino]methyl}$ benzamide;$

 $\label{eq:continuous} $$ $$ $$ V-[2-amino-5-(2-thienyl)phenyl]-4-{[(4-pyridin-3-ylpyrimidin-2-yl)amino]methyl}$ benzamide; $$$

N-[2-amino-5-(2-thienyl)phenyl]-4-[((6-[2-(dimethylamino)ethoxy]-l/-l-benzimidazol-2-yl}thio)methyl]benzamide;

/V-[2-amino-5-(2-thienyl)phenyl]-4-{[(5-chloro-6-fluoro-IH-benzimidazol-2-yl)amino]methyl}benzamide;

N-[2"amino-5-(2-thienyl)phenyl]-5-{[(3,4,5-trimethoxyphenyl)amino]methyl}-l-benzofuran-2-carboxamide;

I\H2-amino-5-(2-thienyl)phenylM-{3,4,5-trimethoxybenzyl)indoline-6-carboxamide;

trans-/V-[2-amino-5-(2-thienyl)phenyl]-3-(4-{[(3,4,5-

trimethoxyphenyl)amino]methyl}phenyl)acrylamide;

N-[2-amino-5-(2-thienyl)phenyl]-4-{[(3-fluoro-4-methoxyphenyl)amino]methyl}benzamide;

N-[2-amino-5-(2-thienyl)phenyl]-4-{[(6-chloro-5-fluoro-l/-/-benzimidazol-2-yl)thio]methyl}benzamide;

and a pharmaceutically acceptable salt of any one or more of the foregoing.

- 33. A compound according to claim 1 for use in inhibting histone deacetylase.
- 34. A compound according to calim 1 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.
- 35. The compound of claim 34, wherein said treatment is effected by inhibiting histone deacetylase,
- 36. The compound of calim 34, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 37. The compound of claim 34, wherein said cell proliferative disease is cancer.
- 38. The compound of claim 37, wherein said cancer is a solid tumor cancer.
- 39. The compound of claim 37, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 40. A pharmaceutical composition comprising a compound according to claim 1 and a pharmaceutically acceptable carrier.
- 41. The pharmaceutical composition of claim 40 further comprising a nucleic acid level inhibitor of histone deacetylase.
- 42. The pharmaceutical composition of claim 41, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 43. The pharmaceutical composition of claim 42, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1 1 SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.

44. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 1.

- 45. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 40.
- 46. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 41.
- 47. The method of claim 45, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 48. The method of claim 45, wherein said cell proliferative disease is cancer.
- 49. The method of claim 48, wherein said cancer is a solid tumor cancer.
- 50. The method of claim 49, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 51. The method of claim 46, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 52. The method of claim 46, wherein said cell proliferative disease is cancer.
- 53. The method of claim 52, wherein said cancer is a solid tumor cancer.
- 54. The method of claim 53, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 55. A compound of the formula

or a pharmaceutically acceptable salt or in vivo hydrolyzable ester or amide thereof, wherein:

- Φ is -NH $_2$ or -OH;
- ring A is a heterocyclyl, wherein if said heterocyclyl contains an -NH- moiety that nitrogen is optionally substituted by a group selected from K;
- R⁵ is a substituent on carbon and is selected from halo, nitro, cyano, hydroxy, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, siilphamoyl, Ci_6-alkyl, C₂₆-alkenyl, C₂-6-alkynyl, CWalkoxy, C₁₋₆-alkanoyl, Ci.₅-alkanoyloxy, *N*-(C₁₋₅-alkyl)amino, N,N-(C₁₋₆-alkyl)₂amino, Ci-₅-alkanoylamino, N-(C₁₋₅-alkyl)carbamoyl, N,N-(C^-alkyl)₂carbamoyl, C₁₋₆-alkylS(O)_a wherein a is O to 2, C₁₋₆-alkoxycarbonyl, *N*-(C₁₋₆-alkyl)sulphamoyl, N,N-(Ci.₆-alkyl)₂sulphamoyl, aryl, aryloxy, arylCi₋₆-alkyl, heterocyclic group, (heterocyclic group)C₁₋₆-alkyl, or a group (B-E-); wherein R⁵, including group (B-E-), is optionally substituted on carbon by one or more W; and wherein if said heterocyclic group contains an -NH- moiety that nitrogen is optionally substituted by J;
- W is halo, nitro, cyano, hydroxy, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C₁₋₆-alkyl, C₂-6-alkenyl, C₂₋₆-alkynyl, Ci.₆-alkoxy, Ci.₆-alkanoyl, Ci.₆-alkanoyloxy, /V-(Ci.₅-alkyl)amino, A/,N-(C₁₋₆-alkyl)₂amino, Ci.₆-alkanoylamino, N-(C₁₋₆-alkyl)carbamoyl, N,N-(C₁₋₅-alkyl)₂carbamoyl, Ci.₅-alkylS(O)_a wherein a is O to 2, Ci.₆-alkoxycarbonyl, N-(Ci.₆-alkyl)sulphamoyl, N,N-(C₁₋₆-alkyl)₂sulphamoyl, or a group (B'-E¹); wherein W, including group (B'-E¹-), is optionally substituted on carbon by one or more Y;
- Y and Z are independently selected from halo, nitro, cyano, hydroxy, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, Ci.₆-alkyl, C₂₋₆-alkenyl, C₂₋₆-alkenyl, C₁₋₆-alkoxy, C₁₋₆-alkanoyl, Ci.₆-alkanoyloxy, N-(Ci.₆-alkyl)amino, N-NiCi-6-alkyl)₂amino, Ci.₆-alkanoylamino, N-(Ci.₆-alkyl)carbamoyl, N,N-(Ci-6-alkyl)₂carbamoyl, C₁₋₆-alkylS(O)_a wherein a is O to 2, Cw-alkoxycarbonyl, N-(C₁₋₆-alkyl)sulphamoyl or N₁N-(Ci₋₆-alkyl)₂sulphamoyl;
- G, J and K are independently selected from C₁₋₈-alkyl, C₁₋₈-alkenyl, C₁₋₈-alkanoyl, Ci₋₈-alkylsulphonyl, C₁₋₈-alkoxycarbonyl, carbamoyl, N-(C₁₋₈-alkyl)carbamoyl, N-NiCi*-alkyOcarbamoyl, benzyloxycarbonyl, benzoyl, phenylsulphonyl, aryl, arylCi₋₆-alkyl or (heterocyclic group)Ci. 6-alkyl; wherein G, J, and K are optionally substituted on carbon by one or more Q; and wherein if said heterocyclic group contains an -NH- moiety that nitrogen is optionally substituted by hydrogen or Ci₆alkyl;

Q is halo, nitro, cyano, hydroxy, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C_{1-6} -alkyl, C_{2} -6-alkenyl, C_{26} -alkynyl, $C_{1.6}$ -alkoxy, d-e-alkanoyl, C_{16} -alkanoyloxy, A-($C_{1.6}$ -alkyl)amino, A-($C_{1.6}$ -alkyl)sulphamoyl, A-A-($C_{1.6}$ -alkyl)amino, A-($C_{1.6}$ -alkyl)sulphamoyl, A-A-($C_{1.6}$ -alkyl)aminoyl, A-A-($C_{1.6}$ -alkyl)aminoyl, aryl, aryloxy, aryl $C_{1.6}$ -alkyl, arylC-alkoxy, heterocyclic group, (heterocyclic group) $C_{1.8}$ -alkyl, (heterocyclic group) $C_{1.6}$ -alkoxy, or a group (B"-E"-); wherein Q, including group (B"-E"-), is optionally substituted on carbon by one or more Z;

- B, B' and B" are independently selected from Ci.₆-alkyl, C₂₋₆-alkenyl, C₂₋₆-alkynyl, C₃₋₈-cycloalkyl, Ca-s-cycloalkylCi-e-alkyl, aryl, arylCi.₆-alkyl, heterocyclic group, (heterocyclic group)C ₁₋₆-alkyl, phenyl or phenylCi.₆-alkyl; wherein B, B' and B" is optionally substituted on carbon by one or more D; and wherein if said heterocyclic group contains an -NH-moiety that nitrogen is optionally substituted by a group selected from G;
- E, E' and E" are independently selected from -N(R^a)-, -0-, -C(O)O-, -OC(O)-, -C(O)-, -N(RT(O)-, -N(R^a)C(O)N(R^b)-, -N(RT(O)O-, -OC(O)N(R^a)-, -C(O)N(R^a)-, S(O)_r, -SO₂N(R³)-, -N(R³JSO₂- wherein R³ and R^b are independently selected from hydrogen or Ci₋₆-alkyl optionally substituted by one or more F and r is 0-2;
- D and F are independently selected from halo, nitro, cyano, hydroxy, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, $Ci._6$ -alkyl, C_{26} -alkenyl, $C_{2\cdot5}$ -alkynyl, $Ci._6$ -alkoxy, $Ci._6$ -alkanoyl, $Ci._6$ -alkanoyloxy, $Ci._6$ -alkyl) $_2$ amino, d-e-alkanoylamino, $Ci._6$ -alkyl) $_2$ amino, d-e-alkanoylamino, $Ci._6$ -alkyl) $_2$ carbamoyl, $Ci._6$ -alkyl $_3$ 0 wherein a is O to 2, d-e-alkoxycarbonyl, MCi-e-alkyUsulphamoyl or N,N-($C_{1\cdot6}$ -alkyl) $_3$ -sulphamoyl;
- m is 0, 1, 2, 3 or 4; wherein the values of \mathbb{R}^5 may be the same or different; \mathbb{R}^6 is halo:
- n is 0, 1 or 2; wherein the values of R^6 are the same or different; and R^1 , R^2 , R^3 , and R^4 are as defined in claim 1.
- 56. The compound of claim 55 wherein:

Ring A is a heterocyclyl;

R⁵ is halo, nitro, cyano, hydroxy, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, Ci.₆-alkyl, C₂₋₆-alkenyl, C₂₋₆-alkynyl, Ci.₆-alkoxy, Ci.

- E is -N(Ra)-, -0-, -C(O)O-, -OC(O)-, -C(O)-, -N(RT(O)-, -C(O)N(Ra)-, -S(0), , -SO₂N(Ra)-, -N(Ra)SO₂- wherein R3 is hydrogen or $C_{1.6}$ -alkyl optionally substituted by one or more D and r is 0-2;
- D is independently selected from halo, nitro, cyano, hydroxy, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C_{1.6}-alkyl, C_{2.6}-alkenyl, C_{2.6}-alkynyl, Ci.₆-alkoxy, C_{1.6}-alkanoyl, Ci-e-alkanoyloxy, MCi-e-alkyDamino, N,ZV-(C_{1.6}-alkyl) ₂amino, C₁.e-alkanoylamino, N-(Ci-e-alkyl)carbamoyl, N,N-(Ci-e-alkyl) ₂carbamoyl, C₁. 6-alkylS(0) _a wherein a is O to 2, Ci.₆-alkoxycarbonyl, N-(Ci-e-alkyl)sulphamoyl and N-N-[Ci-e-alkyl) ₂sulphamoyl;
- G is selected from C₁₄-alkyl, C₁₄-alkanoyl, C_{1,4}-alkylsulphonyl, C^-alkoxycarbonyl, carbamoyl, A/-(Ci₄-alkyl)carbamoyl, A/-(Ci₄-alkyl)carbamoyl, benzyl, benzyloxycarbonyl, benzoyl and phenylsulphonyl;
- m is 0, 1, 2, 3 or 4; wherein the values of R5 are the same or different;

R5 is halo; and

- n is 0, 1 or 2; wherein the values of R6 are the same or different.
- 57. The compound of claim 56 wherein:
 - Ring A is a pyridyl, quinolyl, indolyl, pyrimidinyl, morpholinyl, piperidinyl, piperazinyl, pyradazinyl, pyrazinyl, thiazolyl, thienopyrimidinyl, thienopyridinyl, purinyl, triazinyl, oxazolyl, pyrazolyl, or furanyl; wherein if Ring A contains an -NH- moiety that nitrogen is optionally substituted by a group selected from K;
 - R5 is a substituent on carbon and is selected from halo, amino, Ci_6-alkyl, Ci-e-alkoxy, AZ-(Ci_6-alkyDamino, aryl, aryloxy, arylC 1.6-alkyl, heterocyclic group, (heterocyclic group)C 1.6-alkyl, or a group (B-E-); wherein R5, including group (B-E-), is optionally substituted on carbon by

one or more W; and wherein if said heterocyclic group contains an -NH- moiety that nitrogen is optionally substituted by J;

- W is hydroxy, mercapto, Ci₋₆-alkyl, C^-alkoxy, N,N-[C_{1·6}-akyl)₂amlno or a group (B'-E'-); wherein W, including group (B'-E'-), is optionally substituted on carbon by one or more Y;
- Y and Z are independently selected from halo, nitro, cyano, hydroxy, C^-alkoxy, N,N-(Ci₆-alkyl)₂amino or Ci.₅-alkanoylamino;
- G, J and K are independently selected from C^-alkyl, C_Z-8-alkenyl, C^-alkanoyl, aryl, arylCW alkyl or (heterocyclic group)Ci-6-alkyl; wherein G, J and K are optionally substituted on carbon by one or more Q; and wherein if said heterocyclic group contains an -NH- moiety that nitrogen is optionally substituted by hydrogen or Ci.6-alkyl;
- Q is cyano, hydroxy, Ci.₆-alkoxy, Ci.₆-alkanoyloxy, Ci.₆-alkoxycarbonyl, Ci.₅-alkoxycarbonylamino, aryl, aryloxy or a group (B"-E"-); wherein Q, including group (B"-E"-), is optionally substituted on carbon by one or more Z;
- B, B' and B" are independently selected from (Walkyl, C₂-6-alkenyl, C₂^-alkynyl, C_{3:8}-cycloalkylC_{1:6}-alkyl, aryl, arylCi_{.6}-alkyl, heterocyclic group, (heterocyclic group)Ci.6-alkyl, phenyl or phenylC^-alkyl; wherein B, B¹ and B" are optionally substituted on carbon by one or more D; and wherein if said heterocyclic group contains an -NH-moiety that nitrogen is optionally substituted by a group selected from G;
- E, E' and E" are independently selected from -N(R^a)-, -0-, -C(O)O-, -OC(O)-, -C(O)-, -N(RT(O)-, -N(R^a)C(O)N(R^a)-, .N(RT(O)O -, -OC(O)N(R^a)-, -C(O)N(R^a)-, -S(0)_r, -SO₂N(R³)-, -N(R^a)SO₂- wherein R^a and R^b are independently selected from hydrogen or Ci.₆-alkyl optionally substituted by one or more F and r is 0-2;

D and F are independently selected from halo, Ci_{-6} -alkoxy or $N,N-(Ci_{-6}$ -alkyl)₂amino;

m is 0, 1, 2, 3 or 4; wherein the values of R5 are the same or different;

R⁶ is fluoro or chlbro; and

- n is 0, 1 or 2, wherein the values of R5 are the same or different;
- 58. The compound of claim 57 wherein:
 - Ring A is pyridin-4-yl, pyridin-3-yl, pyriclin-2-yl, quinolin-8-yl, pyrimidin-6-yl, pyrimidin-5-yl, pyrimidin-4-yl, morpholin-4-yl, piperidin-4-yl, piperidin-3-yl, piperdin-2-yl, piperazin-4-yl, pyridazin-5-yl, pyrazin-6-yl, thiazol-2-yl, thien-2-yl, thieno[3,2d]pyrimidinyl,

thieno[3,2b]pyrimidinyl. thieno[3,2b]pyridinyl, purin-6-yl or triazin-6-yl; wherein if Ring A contains an -NH- moiety that nitrogen is optionally substituted by a group selected from K;

- R⁵ is a substituent on carbon and is selected from fluoro, chloro, amino, methyl, ethyl, propyl, methoxy, N-methylamino, N-ethylamino, N-propylamino, N-butylamino, phenyl, naphthylethyl, piperazin-1-yl, piperidin-1-yl, piperidin-4-yl, 2-(thiomethyl)-pyrimidin-4-yl, tetrahydrofuran-2-ylmethyl, tetrahydropyran-2-ylmethyl, I,2,5-thiadiazol-3-ylethyl, piperidin-1-ylmethyl, pyridin-2-ylmethyl, or a group (B-B-); wherein R⁵, including group (B-B-), is optionally substituted on carbon by one or more W; and wherein if said heterocyclic group contains an -NH- moiety that nitrogen is optionally substituted by J;
- W is hydroxy, methyl, ethoxy, IV,N-(diethyl)amino, N,N-(dibutyl)amino, or a group (B'-E'-); wherein W, including group (B'-E'-), is optionally substituted on carbon by one or more Y; Y and Z are independently selected from fluoro, chloro, bromo, nitro, cyano, hydroxy, methoxy, N,N-(dimethyl)amino or methylcarbonylamino;
- G, J and K are independently selected from methyl, ethyl, propyl, pentyl, 2-methylbutyl, butyl, acetyl, benzyl, 3-(pyrrol-l-yl)propyl or pyrrolidin-2-one-(5S)-methyl; wherein G, J and K are optionally substituted on carbon by one or more Q; and wherein if said heterocyclic group contains an -NH- moiety that nitrogen is optionally substituted by hydrogen or methyl;
- Q is cyano, hydroxy, methoxy, ethoxy, methylcarbonyloxy, methoxycarbonyl, *t*-butoxycarbonylamino, phenyl or a group (B"-E"-); wherein Q, including group (B"-E"-), is optionally substituted on carbon by one or more Z;
- B, B' and B" are independently selected from methyl, ethyl, propyl, cyclohexyl, phenyl, benzyl, 1,2,3,4-tetrahydroquinolinyl, 3-morpholinopropyl, 2-morpholinoethyl, 2-pyrrolidin-l-ylethyl, 3-morpholinopropyl, 3-(4-methylpiperazin-l-yl)propyl, 2-piperidin-l-ylethyl, 3-piperidin-l-ylpropyl, pyridin-3-ylmethyl or imidazol-1-ylpropyl; wherein B, B' and B" are optionally substituted on carbon by one or more D; and wherein if said heterocyclic group contains an -NH- moiety that nitrogen is optionally substituted by a group selected from G;
- E, E' and E" are independently selected from -N(R^a)-, -0-, -C(O)-, -NHC(O)-, -N(RT(O)O-; wherein Ra is hydrogen or methyl optionally substituted by one or more F;

D and F are independently selected from fluoro, methoxy or ethoxy;

m is 0, 1, or 2; wherein the values of R⁵ are the same or different;

R6 is fluoro; and

n is O or 1.

59. The compound of claim 55 that is selected from one of the compounds from Tables 1-8 and 13 of WO 03/087057 modified by replacing the terminal moiety:

NH₂ With
$$R^4 \Phi$$
 , wherein Φ , R^1 , R^2 , R^3 , and R^4 are as defined in claim 1.

- 60. A compound according to claim 55 for use in inhibting histone deacetylase.
- 61. A compound according to calim 55 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.
- 62. The compound of claim 61, wherein said treatment is effected by inhibiting histone deacetylase.
- 63. The compound of calim 61, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 64. The compound of claim 61, wherein said cell proliferative disease is cancer.
- 65. The compound of claim 64, wherein said cancer is a solid tumor cancer.
- 66. The compound of claim 64, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 67. A pharmaceutical composition comprising a compound according to claim 55 and a pharmaceutically acceptable carrier.
- 68. The pharmaceutical composition of claim 67 further comprising a nucleic acid level inhibitor of histone deacetylase.
- 69. The pharmaceutical composition of claim 68, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 70. The pharmaceutical composition of claim 69, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1 1 SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.
- 71. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 55.

72. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 67.

- 73. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 68.
- 74. The method of claim 72, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 75. The method of claim 72, wherein said cell proliferative disease is cancer.
- 76. The method of claim 75, wherein said cancer is a solid tumor cancer.
- 77. The method of claim 76, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 78. The method of claim 73, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 79. The method of claim 73, wherein said cell proliferative disease is cancer.
- 80. The method of claim 77, wherein said cancer is a solid tumor cancer.
- 81. The method of claim 78, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 82. A compound of the formula:

the N-oxide forms, the pharmaceutically acceptable addition salts or the stereo-chemically isomeric forms thereof, wherein

 Φ is -NH $_2$ or -OH;

n is 0,1, 2 or 3, wherein when n is 0 then a direct bond is intended;

t is 0, 1, 2, 3 or 4, wherein when t is 0 then a direct bond is intended;

Q, X, Y, and Z are independently N or CH;

R1 is H or as defined in claim 1;

R2, R3, and R4 are as defined in claim 1;

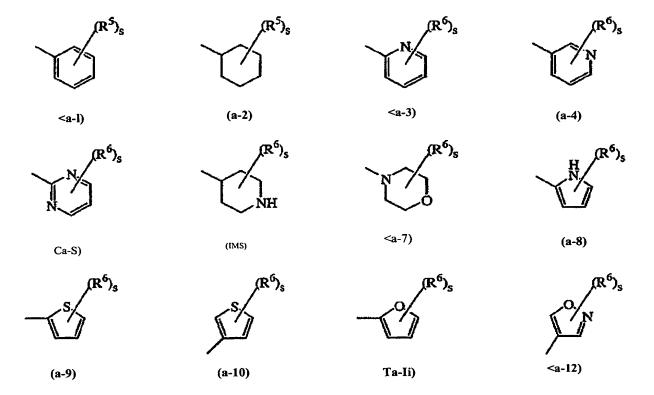
R¹² is hydrogen, halo, hydroxy, amino, nitro, Ci.₆-alkyl, Ci-6-alkyloxy, trifluoromethyl, di(Ci.₆-alkyOamino, hydroxyamino and naphthalenylsulfonylpyrazinyl;

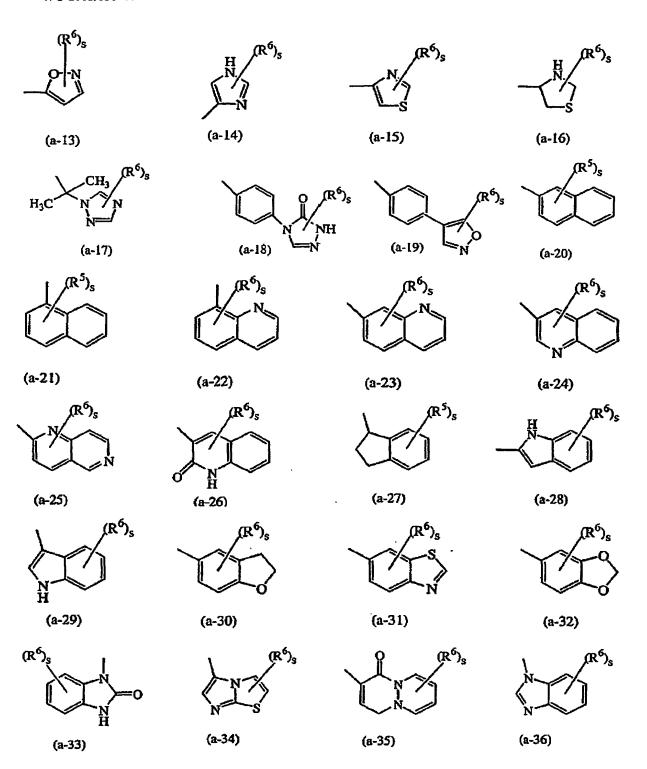
-L- is a direct bond or a bivalent radical selected from Ci₆-alkanediyl, amino, carbonyl and aminocarbonyl;

each R13 is a hydrogen atom, wherein when t is 2, 3, or 4 one of the R13 is optionally aryl;

R¹⁴ is hydrogen, hydroxy, amino, hydroxyC^-alkyl, CWalkyl, Ci₋₆-alkyloxy,arylC₁₋₆-alkyl_F aminocarbonyl, hydroxycarbonyl, aminoCi-₆-alkyl, aminocarbonylCw-alkyl, hydroxycarbonylCi-6-alkyl, hydroxyaminocarbonyl, Ci₋₆-alkyloxycarbo nyl, Ci-s-alkylaminoCi. 6-alkyl or di(C₁₋₆-alkyl)aminoC₁₋₆-alkyl;

Ring A is selected from





$$(a-41)$$
 $(a-42)$ $(a-43)$ $(a-44)$ $(a-45)$ $(a-46)$ $(a-46)$ $(a-47)$ $(a-48)$ $(a-49)$ $(a-50)$ $(a-50)$ $(a-6)$ $(a-6)$

wherein each s is independently 0, 1, 2, 3, 4 or 5;

R5 and R6 are independently selected from hydrogen; halo; hydroxy; amino; nitro; trihaloC $_{1.6}$ -alkyl; trihaloC $_{1.6}$ -alkyloxy; C $_{1.6}$ -alkyl; C $_{1.6}$ -alkyl substituted with aryl and C $_{3.10}$ cycloalkyl; C $_{1.6}$ -alkyloxy; C $_{1.6}$ -alkyloxy; C $_{1.6}$ -alkyloxy; C $_{1.6}$ -alkyloxy; C $_{1.6}$ -alkyloxycarbonyl; C $_{1.6}$ -alkyloxycarbonyl; C $_{1.6}$ -alkyloxycarbonyl; C $_{1.6}$ -alkyloxycarbonyl; cyanoC $_{1.6}$ -alkyl; hydroxyC $_{1.6}$ -alkyloxy; hydroxyC $_{1.6}$ -alkyl)amino; aminoC $_{1.6}$ -alkyloxy; di(C $_{1.6}$ -alkyl)aminocarbonyl; di(hydroxyC $_{1.6}$ -alkyl)amino; (aryl)(C $_{1.6}$ -alkyl)amino; di(C $_{1.6}$ -alkyl)aminoC $_{1.6}$ -alkyl)aminoC $_{1.6}$ -alkyl)aminoC $_{1.6}$ -alkyl)aminoC $_{1.6}$ -alkyl)aminoC $_{1.6}$ -alkyl; arylsulfonyl; arylsulfonylamino; aryloxy; aryloxyC $_{1.6}$ -alkyl; arylC $_{2.6}$ -alkenediyl; di(C $_{1.6}$ -alkyl)amino; di(C $_{1.6}$ -alkyl)aminoC $_{1.6}$ -alkyl)amino; di(C $_{1.6}$ -alkyl)amino; di(C $_{1.6}$ -alkyl)aminoC $_{1.6}$ -alkyl)amino; di(C $_{1.6}$ -alkyl)amino; di(C $_{1.6}$ -alkyl)amino(C $_{1.6}$ -alkyl)amino; di(C $_{1.6}$ -alkyl)amino; di(C $_{1.6}$ -alkyl)amino(C $_{1.6}$ -alkyl)amino(C $_{1.6}$ -alkyl)amino; di(C $_{1.6}$ -alkyl)amino(C $_{1.6}$

 $alkyl)aminoC_{1:6}-alkyl(C_{1:6}-alkyl)amino; \ di(Ci._{6}-alkyl)aminoC_{1:6}-alkyl(C_{1:6}-alkyl)aminoC_{1:6}-alkyl;\\$ $aminosulfonylaminol Ci.e-alky Daminoj \ aminosulfonylaminol Ci.e-alky Damino Ci.e-alky Da$ alkyl)aminosulfonylamino(Ci. s-alkyl)amino; di(Ci.s-alkyl)aminosulfonylamino(Ci. salkyl)aminoCi. 6-alkyl; cyano; thiophenyl; thiophenyl substituted with di(Ci. 6-alkyl)aminoCi. 6alkyKCi-e-alkyDaminoCi-e-alkyI, di(C₁₋₆-alkyI)aminoC ₁₋₆-alkyI, Ci-e-alkyIpiperazinylCi-e-alkyI, hydroxyCi.e-alkylpiperazinylCi.e-alkyl^hydroxyCi.e-alkylpiperazinylCi-e-alkyl, di(Ci. 6-alkyl)aminosulfonylpiperazinylC 16-alkyl, Ci-6-alkyloxypiperidinyl, Ci.6alkyloxypiperidinylCWalkyl, morpholinylCi. 6-alkyl, hydroxyCi.6-alkyl(Ci.6-alkyl)aminoC₁₆alkyl, or di(hydroxyCi. 5-alkyl)aminoCi. 6-alkyl; furanyl; furanyl substituted with hydroxyCi. 6alkyl; benzofuranyl; imidazolyl; oxazolyl; oxazolyl substituted with aryl and Ci s-alkyl; $\hat{C}_{1:6}$ alkyltriazolyl; tetrazolyl; pyrrolidinyl; pyrrolyl; piperidinylCi. s-alkyloxy; morpholinyl; C1.6alkylmorpholinylj morpholinylCi-e-alkyloxy; morpholinylC 1.6-alkyl; morpholinylCi-e-alkylamino; morpholinylC^e-alkylaminoCi-e-alkyl; piperazinyl; Ci.s-alkylpiperazinyl; Ci.s-alkylpiperazinyl; Ci.s-alkylpiperazinyl s-alkyloxy; piperazinylC 1.s-alkyl; naphthaienylsulfonylpiperazinyl; naphthalenylsulfonylpiperidinyl; naphthalenylsulfonyl; Ci-e-alkylpiperazinylCi-e-alkyl; Ci.6alkylpiperazinylCi-e-alkylamino; Ci-e-alkylpiperazinylCi.e-alkylaminoCx.e-alkyl; Ci_salkylpiperazinylsulfonyl; aminosulfonylpiperazinylCi-e-alkyloxy; aminosulfonylpiperazinyl; aminosulfonylpiperazinylCi-e-alkyl; di(C1.6-alkyl)aminosulfonylpiperazinyl; di(Ci.6alkyDaminosulfonylpJperazinylCi-e-alkyl; hydroxyCi ₋₆-alkylpiperazinyl; hydroxyCi-₆alkylpiperazinylCi-6-alkyl; Ci-5-alkyloxyperidinyl; Ci-e-alkyloxypiperidinylCx-6-alkyl; piperidinylaminoC 1,6-alkylamino; piperidinylaminoCi.6-alkylaminoCi.6-alkyl; (Ci.6alkylpiperidinyl)(hydroxyCi_6-alkyl)aminoC 1.6-alkylpiperidinyl)(hydroxyCi_6-alkylpiperidinyl) alkyl)aminoCi. 5-alkylaminoCi-6-alkyl; hydroxyCi-e-alkyloxyCi-e-alkylpiperazinyl; hydroxyCi. 6alkyloxyCi-e-alkylpiperazinylCi-e-alkyl; (hydroxyCi. 6-alkyl)(Ci.6-alkyl)amino; (hydroxyCi.5alky!)(Ci_s-alky!)aminoCi_6-alkyl; hydroxyC^-alkylaminoCi-e-alkyl; di(hydroxyCi. 6alkyl)aminoCi. g-alkyl; pyrrolidinylCi. g-alkyl; pyrrolidinylCi. g-alkyloxy; pyrazolyl; thiopyrazolyl; pyrazolyl substituted with two substituents selected from Ci-6-alkyl and trihaloCi. 6-alkyl; pyridinyl; pyridinyl substituted with Ci. g-alkyloxy, aryloxy or aryl; pyrimidinyl; tetrahydropyrimidinylpiperazinylj tetrahydropyrimidinylpiperazinylCi-e-alkyl; quinolinyl; indolyl; phenyl; phenyl substituted with one, two or three substituents independently selected from halo, amino, nitroA-e-alkyl, Ci. g-alkyloxy, hydroxyCi 4-alkyl, trifluoromethyl,

trifluoromethyloxy, hydroxyCi 4-alkyloxy, Ci4-alkyloxyC14-alkyloxyC14-alkyloxy, Ci4alkyioxycarbonyl,aminoCi 4-alkyloxy, di(Ci4-alkyl)aminoC 14-alkyloxy, di(C14-alkyl)amino, di(C₁₄-alkyl)aminocarbonyl, di(Ci₄-alkyl)aminoCi₄-alkyl, di(C₁₄-alkyl)aminoC₁₄-alkylaminoCi. 4-alkyl, di(Ci4-alkyl)amino(Ci4-alkyl)amino, di(C14-alkyl)amino(C14-alkyl)aminoC14-alkyl, di(C15 ₄-alkyl)aminoC ₁₄-alkyl(Ci₄-alkyl)amino, di(C ₁₄-alkyl)aminoC ₁₄-alkyl(C ₁₄-alkyl)aminoC ₁₄-alkyl, aminosulfonylamino(Ci ₄-alkyl)amino, aminosulfonylamino(C ₁₄-alkyl)aminoC₁₄-alkyl, di(C₁₄alkyl)aminosulfonylamino(C 4-alkyl)amino, di(C 4-alkyl)aminosulfonylamino(Ci 4alkyl)aminoC 4 -alkyl, cyano, piperidinylC14 -alkyloxy, pyrrolidinylC 4 -alkyloxy, aminosulfonylpiperazinyl, aminosulfonylpiperazinylCw-alkyl, di(C14alkyDaminosulfonylpiperazinyl, di(C14-alkyl)aminosulfonylpiperazinylC14-alkyl, hydroxyC14alkylpiperazinyl, hydroxyC 14-alkylpiperazinylC 14-alkyl, Ci4-alkyloxypiperidinyl, Ci4alkyloxypiperdinylC 14-alkyl, hydroxyC14-alkyloxyCw-alkylpiperazinyl^ydroxyCw-alkyloxyCi. ₄-alkylpiperazinylCi ₄-alkyl, (hydroxyCi₄-alkyl)(Ci₄-alkyl)amino, (hydroxyCi₄-alkyl)(Ci₄alkyl)aminoCi ₄ -alkyl, di(hydroxyCi ₄ -atkyl)amino, di(hydroxyC ₁₄ -alkyl)aminoCi ₄ -alkyl, furanyl, furanyl substituted with-CH=CH-CH=CH-, pyrrolidinylCu-alkyl, pyrrolidinylCi 4-alkyloxy, morpholinyl, morpholinylC 4-alkyloxy, morpholiny)Ci 4-alkyl,morpholinylC 4-alkylamino, morpholinylCu-alkylaminoCi- ₄-alkyl, piperazinyl, Ci₄-alkylpiperazinyl, C₁₄-alkylpiperazinylC₁. ₄-alkyloxy, piperazinylC ₁₄-alkyl, Ci₄-alkylpiperazinylC ₁₄-alkyl, C₁₄-alkylpiperazinylCi ₄alkylamino, Ci^.-alkylpiperazinylCi-a-alkylaminoCi^-alkyl, tetrahydropyrimidinylpiperazinyl, tetrahydropyrimidinylpiperazinylCi- a-alkyl, piperidinylaminoCw-alkylamino, piperidinylaminoCu-alkylaminoCu-alkyl, (Ci₄-alkylpiperidiny!)(hydroxyCi₄-alkyl)aminoCi₄alkylamino, (C₁₄-alkylpiperidinyl)(hydroxyCi₄-alkyl)aminoCi₄-alkylaminoCi₄-alkyl, pyridinylCi _a-alkyloxy, hydroxyCi _a-alkylamino, hydroxyCi _a-alkylaminoCi _a-alkyl, di(Ci_aalkyl)aminoCi ₄ -alkylamino, aminothiadiazolyl,aminosulfonylpiperazinylCi ₄ -alkyloxy, and thiophenylCu-alkylamino; the central moiety

$$-N$$
 Z

is optionally bridged (i.e., forming a bicyclic moiety) with a methylene, ethylene or propylene bridge;

each R⁶ and R⁶ can be placed on the nitrogen in replacement of the hydrogen;

aryl in the above is phenyl, or phenyl substituted with one or more substituents each independently selected from halo, Ci_6-alkyi, (-Walkyloxy, trifluoromethyl, cyano, and hydroxycarbonyl.

83. The compound of claim 82 wherein:

n is 1 or 2;

t is 0, 1 or 2;

each Z is nitrogen;

R¹² is hydrogen, nitro, C₁₋₆-alkyloxy, trifluoromethyl, di(C₁₋₅-alkyl) amino, hydroxyamino or naphtalenylsulfonylpyrazinyl;

-L-is a direct bond or a bivalent radical selected from Cw-alkanediyl, carbonyl and aminocarbonyl;

each R13 is hydrogen;

R¹⁴ is hydrogen, hydroxyC _{1.6}-alkyl, aminocarbonyl, hydroxyaminocarbonyl or di(Ci_{.6}-alkyl) aminoCi-s-alkyl;

the A ring is a radical selected from (a-1), (a-7), (a-9),(a-10), (a-12), (a-14), (a-19), (a-20), (a-21), (a-22), (a-23), (a-30), (a-34), (a-49) and (a-50);

each s is independently 0,1, 2 or 5;

each R⁶ are independently selected from hydrogen; halo; nitro; trihaloCi. ₆-alkyl; trihaloCi- ₆-alkyloxy; Ci.₆-alkyl; CWalkyloxy; Ci.₆-alkylsulfonyl; (aryl)(Ci.₅-alkyl)amino; arylsulfonyl; aryloxy; arylC^e-alkenediyl; di(C_{1:6}-alkyl)amino; thiophenyl; thiophenyl substituted with dkCi-e-alkyDaminoCi-e-alkykCi-e-alkyDaminoCi-e-alkyl, dkCi-e-alkyOaminoCi-e-alkyl.Ci-e-alkylpiperazinylCi-e-alkyl, hydroxyCi-₆-alkylpiperazinylCi-e-alkyl, hydroxyCi.₆-alkyloxyCi.₆-alkylpiperazinylCi-e-alkyl, ditCi-e-alkyDaminosulfonylpiperazinylCi-e-alkyl, Ci.₅-alkyloxypiperidinylCi-₆-alkyl, morpholinylCi.₆-alkyl, hydroxyCi-s-alkyKCi-e-alkyDaminoCi-s-alkyl, or difhydroxyCi-e-alkyDaminoCi-₆-alkyl; furanyl; oxazolyl; pyrrolyl; pyrazolyl; pyridinyl; pyridinyl substituted with C_{1.6}-alkyloxy; quinolinyl; indolyl; phenyl; and phenyl substituted with one, two or three substituents independently selected from halo, amino, C_{1.6}-alkyl, C₁-alkyloxy, hydroxyC₁-alkyl, trifluoromethyl, trifluoromethyloxy, di(Ci.₄-alkyl)aminoCi.₄-alkyl, dkCu-alkyDaminoCw-alkyKCu-alkyDaminoC, dilC²-alkylDaminoCw-alkyKCu-alkyDaminoC 1₄-alkyl, hydroxyC 1₄-alkyl, hydroxyC 1₄-alkyl, hydroxyC 1₄-alkyl, di (hydroxyC 1₄-alkyl, hydroxyC 1₄-alkyl, di (hydroxyC 1₄-alkyl, hydroxyC 1₄-alkyl, di (hydroxyC 1₄-alkyl, di (hydroxyC 1₄-alkyl)andinoCw-alkylpiperazinylCw-alkyl, di (hydroxyC 1₄-alkylpiperazinylCw-alkyl), di (hydroxyC 1₄-alkylpiperazinylCw-alkyl, di (hydroxyC 1₄-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinylCw-alkylpiperazinyl

alkyl)aminoCi ₄-alkyl, pyrrolidinylCw-alkyl, pyrrolidinylC _{1.4}-alkyloxy, morpholinylCu-alkyloxy, morpholinylCw-alkyl, and Cw-alkylpiperazinylCw-alkyl, and

the central moiety

 $-\sqrt{-(CH_2)_n}$

is optionally bridged (i.e., forming a bicyclic moiety) with a methylene bridge.

84. The compound of claim 83 wherein:

t is O or 2;

R12 is hydrogen;

-L-is a direct bond;

R¹⁴ is hydrogen;

the A ring is a radical selected from (a-1), (a-9), (a-19), (a-20), (a-21), (a-22), (a-23), (a-49) and (a-50); and

each R⁵ and R⁶ is independently selected from hydrogen; halo; trihaloCi ₆-alkyl; trihaloCi ₆-alkyloxy; Ci ₆-alkyl; C_{1.6}-alkyloxy; arylC^-alkenediyl; di(Ci ₆-alky)amino; thiophenyl; thiophenyl substituted with dKCi^-alkyDaminoCi-s-alkyKC^alkyDaminoCi-ralkyl, di(Ci ₆-alkyDaminoCi-ralkyl, Ci-e-alkylpiperazinylCi-s-alkyl, hydroxyCi-e-alkylpiperazinylCi^-alkyl, hydroxyCi-e-alkylpiperazinylCi-e-alkyl, Ci^-alkyloxypiperidinylC _{1.6}-alkyl, morpholinylCi-6-alkyl, hydroxyCi^-alkyl(Ci^-alkyl)aminoCi-s-alkyl, or di(hydroxyC _{1.6}-alkyDaminoCi-6-alkyl; furanyl; oxazolyl; pyrazolyl; pyridinyl; pyridinyl substituted with Ci ₆-alkyloxy; quinolinyl; indolyl; phenyl; and phenyl substituted with one, two or three substituents independently selected from halo, amino, CWalkyl, C_{1.6}-alkyioxy, hydroxyC _{1.4}-alkyl, trifluoromethyl, trifluoromethyloxy, dKCu-alkyDaminoC^-alkyloxy, di(Ci ₄-alkyl)amino, di(C_{1.4}-alkyl)aminoC_{1.4}-alkyl, di(Ci ₄-alkyl)aminoC _{1.4}-alkyl)aminoCi ₄-alkyl, hydroxyCw-alkyloxyC _{1.4}-alkylpiperazinylCw-alkyl, di(hydroxyC _{1.4}-alkyl)aminoCi ₄-alkyl, pyrrolidinylCu-alkyl pyrrolidinylC^-alkyloxy, morpholinylCWalkyloxy, morpholinylCi ₄-alkyl, and C^-alkylpiperazinylCw-alkyl.

85. The compound of claim 83 wherein:

n is 1;

t is O;

R12 is hydrogen;

-L-is a direct bond;

R14 is hydrogen;

the A ring is a radical selected from (a-1) and (a-20);

each s is independently Oor 1; and

- each R⁶ and R⁶ is independently selected from hydrogen; thiophenyl; thiophenyl substituted with di(C_{1·5}-alkyl)aminoC_{1·6}-alkyl or C^-alkylpiperazinylCi-e-alkyl; furanyl; phenyl; and phenyl substituted with one substituents independently selected from dKCu-alkyOaminoCpr alkyloxy, dKCw-alkyOamino, di(Ci4-alkyl)aminoCi4-alkyl, dilCM-alkyDaminoCw-alkyKCw-alkyl)aminoCi₄-alkyl, pyrrolidinylCWalkyl, pyrrolidinylC^-alkyloxy and Ci₄-alkylpiperazinylCu-alkyl.
- 86. The compound of claim 82 wherein L is a direct bond and R¹² is H.
- 87. The compound of claim 82 wherein:

t is o;

- R¹² is hydrogen, halo, hydroxy, amino, nitro, Ci.₆-alkyl, C_{1.6}-alkyloxy, trifluoromethyl or dKCW alkyOamino;
- -L- is a direct bond or a bivalent radical selected from C_{1.6}-alkanediyl, amino, and carbonyl;
- R¹⁴ is hydrogen, hydroxy, amino, hydroxyCi.₆-alkyl, CWalkyl, C^-alkyloxy, arylCWalkyl, aminoCarbonyl, aminoCi.₆-alkyl, Ci-e-alkylaminoCi-e-alkyl or di(Ci.₆-alkyl)aminoC₁₋₆-alkyl;
- the A ring is a radical selected from (a-1), (a-3), (a-4), (a-5), (a-6), (a-7), (a-8), (a-9), (a-10), can II), (a-12), (a-13), (a-14), (a-15), (a-16), (a-17), (a-18), (a-19), (a-20), (a-21), (a-22), (a-23), (a-24), (a-25), (a-26), (a-28), (a-29), (a-30), (a-31), (a-32), (a-33), (a-34), (a-35), (a-36), (a-37), (a-38), (a-39), (a-40), (a-41), (a-42), (a-44), (a-45), (a-46), (a-47), (a-48) and (a-51); each s is independently 0, 1, 2, 3 or 4;
- R⁵ is hydrogen; halo; hydroxy; amino; nitro; trihaloCi. ₆-alkyl; trihaloCi. ₆-alkyloxy; Ci.₆-alkyl; Ci.₆-alkyloxy; Ci.₆-alkylcarbonyl; C_{1.6}-alkyloxycarbonyl; Ci.₆-alkylsulfonyl; hydroxyCi.₆-alkyl; aryloxy; di(C_{1.6}-alkyl)amino; cyano; thiophenyl; furanyl; furanyl substituted with hydroxyCi.₆-alkyl; benzofuranyl; imidazolyl; oxazolyl; oxazolyl substituted with aryl and Ci.₆-alkyl; C_{1.6}-alkyltriazolyl; tetrazolyl; pyrrolidinyl; pyrrolyl; morpholinyl; Ci.₆-alkylmorpholinyl; piperazinyl; CWalkylpiperazinyl; hydroxyCi.₆-alkylpiperazinyl; Ci.₆-alkyloxypiperidinyl; pyrazoly; pyrazolyl substituted with one or two substituents selected from Ci.₆-alky and trihaloCi-₆-alkyl; pyridinyl; pyridinyl substituted with d ₆-alkyloxy, aryloxy or aryl;

pyrimidinyl; quinolinyl; indole; phenyl; or phenyl substituted with one or two substituents independently selected from halo, Ci_6-alkyl, Ci.6-alkyloxy, or trifluoromethyl;

R6 is hydrogen; halo; hydroxy; amino; nitro; triha!oCi _6-alkyl; trihaloCi _6-alkyloxy; C _1.6-alkyl; Ci. 6-alkyloxy; C _1.6-alkylcarbonyl; CWalkyloxycarbonyl; Ci _6-alkylsulfonyl; hydroxyCi _6-alkyl; aryloxy; di(Ci _6-alkyl)amino; cyano; pyridinyl; phenyl; or phenyl substituted with one or two substituents independently selected from halo, C _1.6-alkyl, Ci. 5-alkyloxy, and trifluoromethyl, and

the central moiety

$$-N$$
 Z

is optionally bridged (i.e., forming a bicyclic moiety) with an ethylene bridge.

88. The compound of claim 82 wherein:

R12 is hydrogen, halo, hydroxy, amino, nitro, Ci.₆-alkyl, CWalkyloxy, trifluoromethyl, hydroxyamino or naphthalenylsulfonylpyrazinyl;

Rw is hydrogen, hydroxy, amino, hydroxyCi. ₆-alkyl, C₁₋₆-alkyloxy, arylCi. ₆-alkyl, aminocarbonyl, hydroxycarbonyl, aminoCi. ₆-alkyl, aminocrbonylCi- ₆-alkyl, hydroxycarbonylCWalkyl, hydroxyaminocarbonyl, Ci-e-alkyloxycarbonyl, Ci.₆-alkylaminoCi. ₆-alkyl or CJi(Ci.₆-alkyl) aminoCi- ₆-alkyl;

the A ring is a radical selected from (a-1), (a-2), (a-3), (a-4), (a-5), (a-6), (a-7), (a-8), (a-9), (a-10), (a-11), (a-12), (a-13), (a-14), (a-15), (a-16), (a-17), (a-18), (a-19), (a-20), (a-21), (a-22), (a-23), (a-24), (a-25), (a-26), (a-27), (a-28), (a-29), (a-30), (a-31), (a-32), (a-33), (a-34), (a-35), (a-36), (a-37), (a-38), (a-39), (a-40), (a-41), (a-42), (a-43) and (a-44); and

each R5 and R6 are independently selected from hydrogen; halo; hydroxy; amino; nitro; trihaloCi _{.6}-alkyl; trihaloCi. ₆-alkyloxy; Ci.₆-alkyl; Ci_{.6}-alkyloxy; Ci.₆-alkyloxy; Ci.₆-alkyloxyCi-e-alkyloxy; Ci.₆-alkyloxyCi. ₅-alkyl; hydroxyCi.₅-alkyl; hydroxyCi-s-alkyloxy; hydroxyCi-₆-alkylamino; aminoC^-alkyloxy; di(C _{1.6}-alkyl)aminocarbonyl; difliydroxyCW alkyDamino; di(Ci-₆-alkyl)aminoCi.₆-alkyloxy; di(Ci.₆-alkyl)aminoCi.₆-alkylamino; arylsulfonyl; arylsulfonylamino; aryloxy; arylC^-alkenediyl; di(C _{1.6}-alkyl)amino; cyano; thiophenyl; thiophenyl substituted with di(Ci ₈-alkyl)aminoCi ₆-alkyl(Ci ₆-alkyl)aminoC ₁₆-alkyt, di(Ci ₈-alkyl)aminoCi-₆-alkyl, Ci-βalkylpiperazinylCi-e-alkyl or di(hydroxyCi-6-alkyl)aminoC _{1.6}-alkyl; furanyl; imidazolyl; Ci.₆-alkyltriazolyl; tetrazolyl; piperidinylCi ₈-alkyloxy; morpholinyl; Ci-₆-alkyloxy; morpholinyl; Ci-₆-alkyl)

alkylmorpholinyl; morpholinylCi. _s-alkyloxy; morpholinylC _{1.s}-alkyl; Ci_{.s}-alkylpiperazinyiCi _{.s}alkyloxy; Ci-e-alkylpiperazinylC^-alkyl; Ci. 6-alkylpiperazinylsulfonyl; aminosulfonylpiperazinylCi-e-alkyloxy; aminosulfonylpiperazinyl; aminosulfonylpiperazinylCi. 6-alkyl; di(Ci, -alkyl)anninosulfonylpiperazinyl; dKCi-e-alkyDaminosulfonylpiperazinylCi-e-alkyl; hydroxyCi-6-alkylpiperazinyl; hydroxyC^-alkylpiperazinylCi-e-alkyl; Ci₋₆-alkyloxyp[peridiny[; Ci-e-alkyloxypiperidinylCi-e-alkyl; hydroxyCi-e-alkyloxyCi-e-alkyipiperazinyl; hydroxyC χ_{e} alkyloxyCi-e-alkylpiperazinylCi-e-alkyl; (hydroxyd-e-alkylXCi-e-alkyDamino; (hydroxyCi. 6alky[)(Ci _-alkyl)aminoCi _-alkyl; pyrrolidinyfC _-alkyloxy; pyrazolyl; thiopyrazolyl; pyrazolyl substituted with two substituents selected from C^e-alky) or trihaloC^-alkyl; pyridinyl; pyridinyl substituted with C1-6-alkyloxy or aryl; pyrimidinyl; quinolinyl; phenyl; and phenyl substituted with one, two or three substituents independently selected from halo, amino, Ci. 6 alkyl, Ci. 6 alkyloxy, hydroxyC 14 alkyl, trifluoromethyl, trifluoromethyloxy, hydroxyC 14 alkoxy, C₁₄-alkyloxyC₁₄-alkoxy, aminoCi₄-alkyloxy, di(Ci₄-alkyl)aminoCi₄-alkyloxy, di(C₁₄alkyDamino, piperidinylC₁₋₄-alkyloxy, pyrrolidinylC₁₄-alkyloxy, aminosulfonylpiperazinyl, aminosulfonylpiperazinylCw-alkyl, di(Ci4-alkyl)aminosulfonylpiperazinyl, di(Ci4alkyl)aminosulfonylpiperazinylC 44-alkyl, hydroxyC 44-alkylpiperazinyl, hydroxyC 44hydroxyCi. alkylpiperazinylC 14 -alkyl, Ci4-alkyloxypiperidinyl, Cu-alkoxypiperidinylC^-alkyl, ₄-alkyloxyC ₁₄-alkylpiperazinyl, hyroxyCw-alkoxyCw-alkylpiperazinylCw-alkyl, hydroxyCi ₄ $alkyl)(C_{4}-alkyl)amino, \ (hydroxyCi_{4}-alkyl)(Ci_{4}-alkyl)aminoGi_{4}-alkyl, \ pyrrolidinylCi_{4}-alkoxy, \ alkyl)(Ci_{4}-alkyl)aminoGi_{4}-alkyl), \ alkyl)(Ci_{4}-alkyl)aminoGi_{4}-alkyl), \ byrrolidinylCi_{4}-alkyl), \ byrrolidinylCi_{4}-a$ morpholinylCi- 4-alkyloxy, morpholinylC 4-alkyl, C4-alkylpiperazinylCi 4-alkoxy, C4alkylpiperazinylCi-4-alkyl, hydroxyCi4-alkylamino, di(hydroxyCi4-alkyl)amino, di(C14alkyl)aminoCi ₄-alkylamino, aminothiadiazolyl, aminosulfonylpiperazinylC^-alkyloxy, and thiophenylCw-alkylamino.

89. The compound of claim 82 that is selected from one of the compounds of pages 21 and 22 and Table F-I of WO 03/076422 wherein the terminal hydroxamic acid moiety (HO-NH-C(O)-) is replaced with

wherein Φ , R^1 , R^2 , R^3 , and R^4 are as defined in claim 1.

- 90. A compound according to claim 82 for use in inhibting histone deacetylase.
- 91. A compound according to calim 82 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.
- 92. The compound of claim 91, wherein said treatment is effected by inhibiting histone deacetylase.
- 93. The compound of calim 91, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 94. The compound of claim 91, wherein said cell proliferative disease is cancer.
- 95. The compound of claim 94, wherein said cancer is a solid tumor cancer.
- 96. The compound of claim 94, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 97. A pharmaceutical composition comprising a compound according to claim 82 and a pharmaceutically acceptable carrier.
- 98. The pharmaceutical composition of claim 97 further comprising a nucleic acid level inhibitor of histone deacetylase.
- 99. The pharmaceutical composition of claim 98, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 100. The pharmaceutical composition of claim 99, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:I, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.
- 101. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 82.
- 102. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 97.
- 103. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising

administering to said individual a treatment effective amount of the pharmaceutical composition of claim 98.

- 104. The method of claim 102, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 105. The method of claim 102, wherein said cell proliferative disease is cancer.
- 106. The method of claim 102, wherein said cancer is a solid tumor cancer.
- 107. The method of claim 106, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 108. The method of claim 103, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 109. The method of claim 103, wherein said cell proliferative disease is cancer.
- 110. The method of claim 109, wherein said cancer is a solid tumor cancer.
- 111. The method of claim 110, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 112. A compound of the formula:

or a pharmaceutically acceptable salt thereof, wherein

 Φ is -NH $_2$ or -OH;

R1 is H or as defined in paragraph claim 1;

R2, R3, and R4 are as defined in paragraph claim 1;

n is 0, 1, 2 or 3 and when n is 0 then a direct bond is intended;

R is selected from the group consisting of hydrogen, halogen, -NH₂, nitro, hydroxy, aryl, heterocyclyl, C_3 - C_8 -cycloalkyl, heteroaryl, C_1 - C_7 -akyl, haloalkyl, C_1 - C_7 -alkenyl, C_1 - C_7 -alkyl-arylsulfanyl, C_1 - C_7 -alkyl-arylsulfinyl, C_1 - C_7 -alkyl-arylsulfinyl, C_1 - C_7 -alkyl-arylaminosulfonyl, C_1 - C_7 -alkyl-arylamine, C_1 - $C_$

- R¹² is hydrogen, halo, hydroxy, amino, nitro, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl, di(C₁₋₆alkyl)amino, hydroxyamino or naphtalenylsulfonylpyrazinyl;
- is hydrogen, C₁₋₆alkyl, arylC₂₋₆alkenediyl, furanylcarbonyl, naphtalenylcarbonyl, -C(O)phenylR⁹, C₁₋₆alkylaminocarbonyl, aminosulfonyl, arylaminosulfonyl, aminosulfonylamino, di(C₁₋₆alkyl)aminosulfonylamino, arylaminosulfonylamino, aminosulfonylaminoC₁₋₆alkyl, di(C₁₋₆alkyl)aminosulfonylaminoC₁₋₆alkyl, arylaminosulfonylaminoC₁₋₆alkyl, di(C₁₋₆alkyl)aminoC₁₋₆alkyl, C₁₋₁₂alkylsulfonyl, di(C₁₋₆alkyl)aminosulfonyl, trihaloC₁₋₆alkylsulfonyl, di(aryl)C₁₋₆alkylcarbonyl, thiophenylC₁₋₆alkylcarbonyl, pyridinylcarbonyl or arylC₁₋₆alkylcarbonyl

wherein each R^9 is independently selected from phenyl; phenyl substituted with one, two or three substituents independently selected from halo, amino, $C_{1.6}$ alkyl, $C_{1.6}$ alkyloxy, hydroxy $C_{1.4}$ alkyl, hydroxy $C_{1.4}$ alkyloxy, amino $C_{1.4}$ alkyloxy, di($C_{1.6}$ alkyl)amino $C_{1.6}$ alkyl, hydroxy $C_{1.4}$ alkylpiperazinyl $C_{1.4}$ alkyl, $C_{1.4}$ alkylpiperazinyl $C_{1.4}$ alkyl, hydroxy $C_{1.4}$ alkyl)amino $C_{1.4}$ alkyl, pyrrolidinyl $C_{1.4}$ alkyloxy, morpholinyl $C_{1.4}$ alkyl)amino $C_{1.4}$ alkyl, pyrrolidinyl $C_{1.4}$ alkyl)amino $C_{1.6}$ alkyl)amino $C_{1.4}$ alkyloxy, $C_{1.4}$ alkylpiperazinyl $C_{1.4}$ alkyl, di(hydroxy $C_{1.4}$ alkyl)amino $C_{1.6}$ alkyl, di(hydroxy $C_{1.4}$ alkyl)amino $C_{1.6}$ alkyl)amino $C_{1.6}$ alkyl, di(hydroxy $C_{1.4}$ alkyl)amino $C_{1.4}$ alkyloxy, $C_{1.4}$ alkylpiperazinyl $C_{1.4}$ alkyl, di(hydroxy $C_{1.4}$ alkyl)amino $C_{1.4}$ alkyl or morpholinyl $C_{1.4}$ alkyloxy.

R¹⁴ is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyloxy, arylC₁₋₆alkyl, aminocarbonyl, hydroxycarbonyl, aminoC₁₋₆alkyl, aminocarbonylC₁₋₆alkyl, hydroxycarbonylC₁₋₆alkyl, hydroxyaminocarbonyl, C₁₋₆alkyloxycarbonyl, C₁₋₆alkylaminoC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl; when R¹³ & R¹⁴ are present on the same carbon atom, R¹³ & R¹⁴ together may form a bivalent radical of formula

-C(O)-NH-CH₂-NR¹⁰- (a-1) wherein R¹⁰ is hydrogen or aryl;

when R¹³ & R¹⁴ are present on adjacent carbon atoms, R¹³ & R¹⁴ together may form a bivalent radical of formula

=CH-CH=CH-CH= (b-1);

aryl in the above is phenyl, or phenyl substituted with one or more substituents each independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl, cyano or hydroxycarbonyl.

113. The compound of claim 112 wherein:

n is 0 or1;

$$Q$$
 is $-CR$, or $-CH$;

R¹² is hydrogen or nitro;

 $\label{eq:calculate_continuous_continuous} $$R^{13}$ is C_{1-6}alkyl, arylC_{2-6}alkenediyl, furanylcarbonyl, naphtalenylcarbonyl, $$C_{1-6}$alkylaminocarbonyl, aminosulfonyl, $$di(C_{1-6}$alkyl)$aminosulfonylamino$$C_{1-6}$alkyl, $$di(C_{1-6}$alkyl)$aminoc$$C_{1-6}$alkyl, C_{1-12}alkylsulfonyl, $$di(C_{1-6}$alkyl)$aminosulfonyl, $$trihalo$$C_{1-6}$alkylsulfonyl, $$di(aryl)$$C_{1-6}$alkylcarbonyl, thiophenyl$$C_{1-6}$alkylcarbonyl, $$$

R¹⁴ is hydrogen;

when R¹³ & R¹⁴ are present on the same carbon atom R¹³ & R¹⁴ together may form a bivalent radical of formula (a-1) wherein R¹⁰ is aryl;

when R¹³ & R¹⁴ are present on adjacent carbon atoms R¹³ & R¹⁴ together may form a bivalent radical of formula (b-1).

114. The compound of claim 112 wherein:

pyridinylcarbonyl or arylC_{1.6}alkylcarbonyl;

n is 1;

$$Q$$
 is $-CR$, $-CR$, or $-CH$;
 Z is nîtrogen;

R¹² is hydrogen;

R¹³ is naphtalenylcarbonyl, C₁₋₁₂alkylsulfonyl or di(aryl)C₁₋₆alkylcarbonyl; R¹⁴ is hydrogen.

- 115. The compound of claim 112 wherein R¹² is H.
- 116. The compound of claim 112 wherein:

R¹² is hydrogen, halo, hydroxy, amino, nitro, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl or di(C₁₋₆alkyl)amino;

- is hydrogen, C₁₋₆alkyl, arylC₂₋₆alkenediyl, furanylcarbonyl, naphtalenylcarbonyl, -C(O)phenylR⁹, C₁₋₆alkylaminocarbonyl, aminosulfonyl, arylaminosulfonyl, aminosulfonylamino, di(C₁₋₆alkyl)aminosulfonylamino, di(C₁₋₆alkyl)aminoC₁₋₆alkyl, C₁₋₁₂alkylsulfonyl, di(C₁₋₆alkyl)aminosulfonyl or pyridinylcarbonyl wherein each R⁹ is independently selected from phenyl; phenyl substituted with one, two or three substituents independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy; or thiophenyl;
- R¹⁴ is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyloxy, arylC₁₋₆alkyl, aminocarbonyl, aminoC₁₋₆alkyl, C₁₋₆alkylaminoC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl.
- 117. The compound of claim 112 wherein:
- R¹² is hydrogen, halo, hydroxy, amino, nitro, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl or di(C₁₋₆alkyl)amino;
- R¹³ is hydrogen, C₁₋₆alkyl, arylC₂₋₆alkenediyl, furanylcarbonyl, naphtalenylcarbonyl, -C(O)phenylR⁹, C₁₋₆alkylaminocarbonyl, aminosulfonyl, arylaminosulfonyl, aminosulfonylamino, di(C₁₋₆alkyl)aminoC₁₋₆alkyl)aminosulfonyl, di(C₁₋₆alkyl)aminosulfonyl or pyridinylcarbonyl wherein each R⁹ is independently selected from phenyl; phenyl substituted with one, two or three substituents independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy; or thiophenyl; and
- R¹⁴ is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyloxy, arylC₁₋₆alkyl, aminocarbonyl, aminoC₁₋₆alkyl, C₁₋₆alkylaminoC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl.
- 118. The compound of claim 112 wherein:

n is 0 or 1; Q is or

-NHC(O)C₁₋₆alkanediylSH; R^{12} is hydrogen or nitro: R^{13} is C_{1-6} alkyl, arylC₂₋₆alkenediyl, furanylcarbonyl, naphtalenylcarbonyl, C_{1-6} alkylaminocarbonyl, aminosulfonyl, di(C_{1-6} alkyl)aminosulfonylaminoC₁₋₆alkyl, di(C_{1-6} alkyl)aminosulfonyl, di(C_{1-6} alkyl)aminosulfonyl, trihaloC₁₋₆alkylsulfonyl, di(aryl)C₁₋₆alkylcarbonyl, thiophenylC₁₋₆alkylcarbonyl, pyridinylcarbonyl or arylC₁₋₆alkylcarbonyl; R^{14} is hydrogen; when R^{13} and R^{14} are present on the same carbon atom R^{13} & R^{14} together may form a bivalent radical of formula (a-1) wherein R^{10} is aryl; or when R^{13} & R^{14} are present on adjacent carbon atoms R^{13} & R^{14} together may formula (b-1).

119. The compound of claim 112 wherein:

n is 1; Q is \subset ; Z is nitrogen; R^{12} is hydrogen; R^{13} is naphthalenylcarbonyl, C_{1-12} alkylsulfonyl or di(aryl) C_{1-6} alkylcarbonyl; and R^{14} is hydrogen.

120. The compound of claim 112 that is selected from one of

wherein the terminal hydroxamic acid moiety (-C(O)-NH-OH) is replaced with

wherein Φ , R^1 , R^2 , R^3 , and R^4 are as defined in accordance with claim 1.

- 121. A compound according to claim 112 for use in inhibting histone deacetylase.
- 122. A compound according to calim 112 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.
- 123. The compound of claim 122, wherein said treatment is effected by inhibiting histone deacetylase.
- 124. The compound of calim 122, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 125. The compound of claim 122, wherein said cell proliferative disease is cancer.
- 126. The compound of claim 125, wherein said cancer is a solid tumor cancer.

127. The compound of claim 125, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.

- 128. A pharmaceutical composition comprising a compound according to claim 112 and a pharmaceutically acceptable carrier.
- 129. The pharmaceutical composition of claim 128 further comprising a nucleic acid level inhibitor of histone deacetylase.
- 130. The pharmaceutical composition of claim 129, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 131 The pharmaceutical composition of claim 130, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID ID IMo:II, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.
- 132. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 112.
- 133. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 128.
- 134. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 129.
- 135. The method of claim 133, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 136. The method of claim 133, wherein said cell proliferative disease is cancer.
- 137. The method of claim 136, wherein said cancer is a solid tumor cancer.

138. The method of claim 137, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.

- 139. The method of claim 134, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 140. The method of claim 134, wherein said cell proliferative disease is cancer.
- 141. The method of claim 140, wherein said cancer is a solid tumor cancer.
- 142. The method of claim 141, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 143. A compound of the formula:

or a pharmaceutically acceptable salt thereof, wherein

 Φ is -NH₂ or -OH;

R1 is H or as defined in claim 1;

R², R³, and R⁴ are as defined in claim 1;

n is 0, 1, 2 or 3 and when n is Q then a direct bond is intended;

m Is 0 or 1 and when m is 0 then a direct bond is intended;

t is 0, 1, 2, 3 or 4 and when t is 0 then a direct bond is intended;

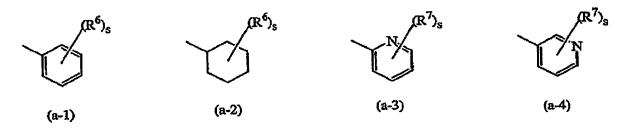
X is nitrogen or -C

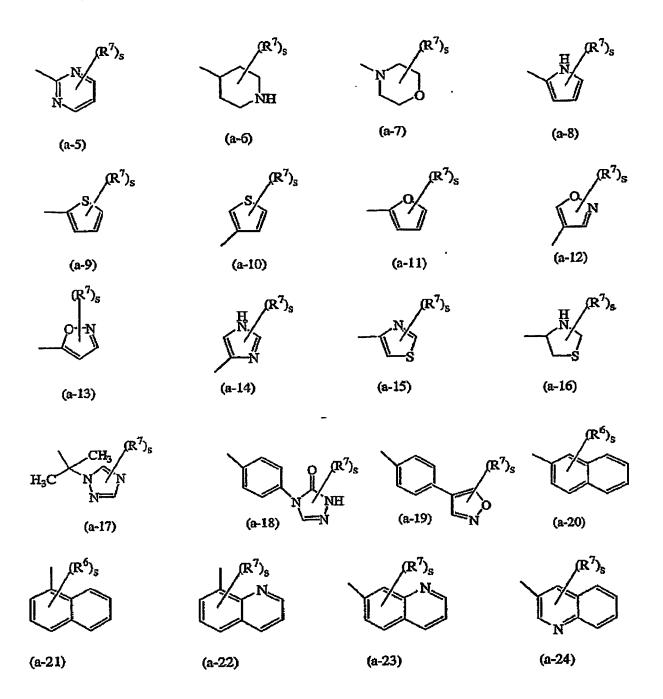
Y is nitrogen or —C

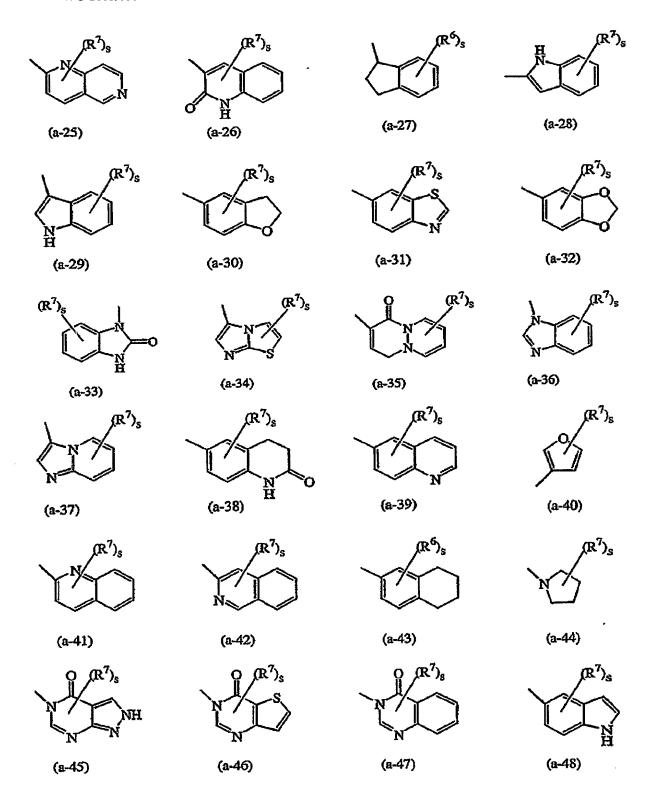
R is selected from the group consisting of hydrogen, halogen, -NH₂, nitro, hydroxy, aryl, heterocyclyl, C₃-C₈-cycloalkyl, heteroaryl, C₁-C₇-akyl, haloalkyl, C₁-C₇-alkenyl, C₁-C₇-alkyl-aryloxy, C₁-C₇-alkyl-arylsulfanyl, C₁-C₇-alkyl-arylsulfinyl, C₁-C₇-alkyl-arylaminosulfonyl, C₁-C₇-alkyl-arylamine, C₁-C₇-alkynyl-C(0)-amine, C₁-C₇-alkenyl-C(0)-amine, C₁-C₇-alkynyl-R⁹, C₁-C₇-alkenyl-R⁹ wherein R⁹ is hydrogen, hydroxy, amino, C₁-C₇-alkyl or C₁-C₇-alkoxy;

- R¹² is hydrogen, halo, hydroxy, amino, nitro, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl, di(C₁₋₆alkyl)amino, hydroxyamino or naphtalenylsulfonylpyrazinyl;
 - -L- is a direct bond or a bivalent radical selected from C₁₋₆alkanediyl, C₁₋₆alkanediyloxy, amino, carbonyl or aminocarbonyl;
- each R¹³ is independently represents a hydrogen atom and one hydrogen atom can be replaced by a substituent selected from aryl;
- is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyloxy, arylC₁₋₆alkyl, aminocarbonyl, hydroxycarbonyl, aminoC₁₋₆alkyl, aminocarbonylC₁₋₆alkyl, hydroxycarbonylC₁₋₆alkyl, hydroxyaminocarbonyl, C₁₋₆alkyloxycarbonyl, C₁₋₆alkylaminoC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl;
- is hydrogen, C₁₋₆alkyl, C₃₋₁₀cycloalkyl, hydroxyC₁₋₆alkyl, C₁₋₆alkyloxyC₁₋₆alkyl, di(C₁₋₆alkyl)aminoC₁₋₆alkyl or aryl;









$$(a-49)$$
 $(a-51)$

wherein each s is independently 0, 1, 2, 3, 4 or 5;

each R⁶ and R⁷ are independently selected from hydrogen; halo; hydroxy; amino; nitro;

 $trihaloC_{1-6}$ alkyl; $trihaloC_{1-6}$ alkyloxy; C_{1-6} alkyl; C_{1-6} alkyl substituted with aryl and

 $C_{3\text{--}10} \\ \text{cycloalkyl}; C_{1\text{--}6} \\ \text{alkyloxy}; C_{1\text{--}6} \\ \text{$

 $C_{1\text{-}6} alkyloxy carbonyl; \ C_{1\text{-}6} alkylsulfonyl; \ cyano C_{1\text{-}6} alkyl; \ hydroxy C_{1\text{-}6} alkyl;$

hydroxyC₁₋₆alkyloxy; hydroxyC₁₋₆alkylamino; aminoC₁₋₆alkyloxy;

 $\label{eq:condition} \mbox{di}(C_{1-6}\mbox{alkyl}) \mbox{amino; (aryl)} (C_{1-6}\mbox{alkyl}) \mbox{amino; (aryl)} (C_{1-6}\mbox{alkyl}) \mbox{amino; }$

 $di(C_{1-6}alkyl)aminoC_{1-6}alkyloxy; di(C_{1-6}alkyl)aminoC_{1-6}alkylamino;$

 $\label{eq:continuous} \mbox{di}(C_{1\text{-}6}\mbox{alkyl}) a \mbox{mino} C_{1\text{-}6}\mbox{alkyl}; \mbox{arylsulfonylamino}; \\ \mbox{arylsulfonylamino};$

aryloxy; aryloxyC₁₋₆alkyl; arylC₂₋₆alkenediyl; di(C₁₋₆alkyl)amino;

 $\label{eq:continuous} \mbox{di}(C_{1\mbox{-}6}\mbox{alkyl}) a \mbox{mino}(C_{1\mbox{-}6}\mbox{alkyl}) a \mbox{mino}(C_{1\mbox{-}6}\mbox{alkyl}) a \mbox{mino};$

 $di(C_{1\text{-}6}alkyl)amino(C_{1\text{-}6}alkyl)aminoC_{1\text{-}6}alkyl;$

 $di(C_{1-6}alkyl)aminoC_{1-6}alkyl(C_{1-6}alkyl)amino;$

 $\label{eq:continuous} \begin{array}{l} \text{di}(C_{1\text{-}6}alkyl) amino C_{1\text{-}6}alkyl (C_{1\text{-}6}alkyl) amino C_{1\text{-}6}alkyl; \end{array}$

aminosulfonylamino(C1-6alkyl)amino;

aminosulfonylamino(C1-6alkyl)aminoC1-6alkyl;

 $di(C_{1\text{-}6}alkyl) amino sulfonylamino (C_{1\text{-}6}alkyl) amino;\\$

 $di(C_{1-6}alkyl)aminosulfonylamino(C_{1-6}alkyl)aminoC_{1-6}alkyl; cyano; thiophenyl;$

thiophenyl substituted with $di(C_{1-6}alkyl)aminoC_{1-6}alkyl(C_{1-6}alkyl)aminoC_{1-6}alkyl$,

 $\label{eq:continuous} \mbox{di}(C_{1\mbox{-}6}\mbox{alkyl}, amino C_{1\mbox{-}6}\mbox{alkyl}, C_{1\mbox{-}6}\mbox{alkylpiperazinyl} C_{1\mbox{-}6}\mbox{alkyl},$

 $hydroxyC_{1-6}alkylpiperazinylC_{1-6}alkyl,$

 $hydroxy C_{1\text{-}6} alkyloxy C_{1\text{-}6} alkylpiperazinyl C_{1\text{-}6} alkyl,$

di(C₁₋₆alkyl)aminosulfonylpiperazinylC₁₋₆alkyl, C₁₋₆alkyloxypiperidinyl, C₁₋₆alkyloxypiperidinylC₁₋₆alkyl, morpholinylC₁₋₆alkyl, hydroxyC₁₋₆alkyl(C₁₋₆alkyl)aminoC₁₋₆alkyl, or di(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkyl; furanyl; furanyl substituted with hydroxyC₁₋₆alkyl; benzofuranyl; imidazolyl; oxazolyl; oxazolyl substituted with aryl and C_{1-6} alkyl; C_{1-6} alkyltriazolyl; tetrazolyl; pyrrolidinyl; pyrrolyl; piperidinylC₁₋₆alkyloxy; morpholinyl; C₁₋₆alkylmorpholinyl; morpholinylC₁₋₆alkyloxy; morpholinylC₁₋₆alkyl; morpholinylC₁₋₆alkylamino; morpholinylC₁₋₆alkylaminoC₁₋₆alkyl; piperazinyl; C₁₋₆alkylpiperazinyl; C₁₋₆alkylpiperazinylC₁₋₆alkyloxy; piperazinylC₁₋₆alkyl; naphtalenylsulfonylpiperazinyl; naphtalenylsulfonylpiperidinyl; naphtalenylsulfonyl: C₁₋₆alkylpiperazinylC₁₋₆alkyl; C₁₋₆alkylpiperazinylC₁₋₆alkylamino; C_{1-6} alkylpiperazinyl C_{1-6} alkylamino C_{1-6} alkyl; C_{1-6} alkylpiperazinylsulfonyl; aminosulfonylpiperazinylC₁₋₆alkyloxy; aminosulfonylpiperazinyl; aminosulfonylpiperazinylC₁₋₆alkyl; di(C₁₋₆alkyl)aminosulfonylpiperazinyl; di(C₁₋₆alkyl)aminosulfonylpiperazinylC₁₋₆alkyl; hydroxyC₁₋₆alkylpiperazinyl; hydroxyC₁₋₆alkylpiperazinylC₁₋₆alkyl; C₁₋₆alkyloxypiperidinyl; C_{1-6} alkyloxypiperidinyl C_{1-6} alkyl; piperidinylamino C_{1-6} alkylamino; piperidinylaminoC₁₋₆alkylaminoC₁₋₆alkyl; (C₁₋₆alkylpiperidinyl)(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkylamino; (C₁₋₆alkylpiperidinyl)(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkylaminoC₁₋₆alkyl; hydroxyC₁₋₆alkyloxyC₁₋₆alkylpiperazinyl; hydroxyC₁₋₆alkyloxyC₁₋₆alkylpiperazinylC₁₋₆alkyl; $(hydroxyC_{1-6}alkyl)(C_{1-6}alkyl)amino; (hydroxyC_{1-6}alkyl)(C_{1-6}alkyl)aminoC_{1-6}alkyl;$ hydroxyC₁₋₆alkylaminoC₁₋₆alkyl; di(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkyl; pyrrolidinylC₁₋₆alkyl; pyrrolidinylC₁₋₆alkyloxy; pyrazolyl; thiopyrazolyl; pyrazolyl substituted with two substituents selected from C_{1-6} alkyl or trihalo C_{1-6} alkyl;

pyridinyl; pyridinyl substituted with C_{1-6} alkyloxy, aryloxy or aryl; pyrimidinyl; tetrahydropyrimidinylpiperazinyl; tetrahydropyrimidinylpiperazinyl C_{1-6} alkyl;

quinolinyl; indole; phenyl; phenyl substituted with one, two or three substituents

independently selected from halo, amino, nitro, C1-6alkyl, C1-6alkyloxy, hydroxyC_{1.4}alkyl, trifluoromethyl, trifluoromethyloxy, hydroxyC_{1.4}alkyloxy, C1_4alkylsulfonyl, C1_4alkyloxyC1_4alkyloxy, C1_4alkyloxycarbonyl, aminoC₁₋₄alkyloxy, di(C1-4alkyl)aminoC1-4alkyloxy, di(C1-4alkyl)amino, di(C1-4alkyl)aminocarbonyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)amino(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(C1-4alkyl)aminoC1-4alkyl(C1-4alkyl)amino, di(C₁₋₄alkyl)aminoC₁₋₄alkyl(C₁₋₄alkyl)aminoC₁₋₄alkyl, aminosulfonylamino(C1-4alkyl)amino, aminosulfonylamino(C1-4alkyl)aminoC1-4alkyl, di(C1_4alkyl)aminosulfonylamino(C1_4alkyl)amino, di(C₁₋₄alkyl)aminosulfonylamino(C₁₋₄alkyl)aminoC₁₋₆alkyl, cyano, piperidinylC₁₋₄alkyloxy, pyrrolidinylC₁₋₄alkyloxy; aminosulfonylpiperazinyl, aminosulfonylpiperazinylC₁₋₄alkyl, di(C₁₋₄alkyl)aminosulfonylpiperazinyl, di(C₁₋₄alkyl)aminosulfonylpiperazinylC₁₋₄alkyl, hydroxyC₁₋₄alkylpiperazinyl, hydroxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, C₁₋₄alkyloxypiperidinyl,

 C_{1-4} ałkyloxypiperidinyl C_{1-4} alkyl, hydroxy C_{1-4} ałkyloxy C_{1-4} alkylpiperazinyl, hydroxy C_{1-4} alkyloxy C_{1-4} alkylpiperazinyl C_{1-4} alkyl,

 $\label{eq:control_problem} \begin{align*} $$ (hydroxyC_{1-4}alkyl)(C_{1-4}alkyl)aminoC_{1-4}alkyl)aminoC_{1-4}alkyl)aminoC_{1-4}alkyl)aminoC_{1-4}alkyl)aminoC_{1-4}alkyl, furanyl, furanyl substituted with -CH=CH-CH=CH-, pyrrolidinylC_{1-4}alkyl, pyrrolidinylC_{1-4}alkyloxy, morpholinylC, morpholinylC_{1-4}alkyloxy, morpholinylC, alkyloxy, morpholinylC, alkyloxy,$

morpholinylC₁₋₄alkylamino, morpholinylC₁₋₄alkylaminoC₁₋₄alkyl, piperazinyl, C₁₋₄alkylpiperazinyl, C₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkyl, C₁₋₄alkylpiperazinylC₁₋₄alkylamino, C₁₋₄alkylpiperazinylC₁₋₄alkylaminoC₁₋₆alkyl, tetrahydropyrimidinylpiperazinyl, tetrahydropyrimidinylpiperazinylC₁₋₄alkyl, piperidinylaminoC₁₋₄alkylamino, piperidinylaminoC₁₋₄alkylaminoC₁₋₄alkyl, (C₁₋₄alkylpiperidinyl)(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkylamino, (C₁₋₄alkylpiperidinyl)(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkylaminoC₁₋₄alkyl, pyridinylC₁₋₄alkyloxy, hydroxyC₁₋₄alkylamino, hydroxyC₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkylamino, aminothiadiazolyl, aminosulfonylpiperazinylC₁₋₄alkyloxy, or thiophenylC₁₋₄alkylamino; each R⁶ and R⁷ can be placed on the nitrogen in replacement of the hydrogen; aryl in the above is phenyl, or phenyl substituted with one or more substituents each independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl, cyano or

hydroxycarbonyl.

```
The compound of claim 143 wherein:
144.
n is 1;
m is 0 or 1:
t is 0.1 or 2;
    R<sup>12</sup> is hydrogen or C<sub>1-6</sub>alkyl;
      -L- is a direct bond;
     R14 is hydrogen;
    R<sup>15</sup> is hydrogen;
                 is a radical selected from (a-1), (a-20), (a-25), (a-27), (a-28), (a-29), (a-41)
      or (a-42);
       each s is independently 0, 1, 2 or 3;
      each R^6 is independently selected from hydrogen, halo, C_{1-6}alkyloxy.
           The compound of claim 143 wherein:
145.
    R^{15}
```

R⁵ is hydrogen, C₁₋₆alkyl, C₃₋₁₀cycloalkyl, hydroxyC₁₋₆alkyl, C₁₋₆alkyloxyC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl;

(a-9), (a-10), (a-11), (a-12), (a-13), (a-14), (a-15), (a-16), (a-17), (a-18), (a-19), (a-20), (a-21), (a-22), (a-23), (a-24), (a-25), (a-26), (a-27), (a-28), (a-29), (a-30), (a-31), (a-32), (a-33), (a-34), (a-35), (a-36), (a-37), (a-38), (a-39), (a-40), (a-41), (a-42) (a-43) or (a-44);

each R⁶ and R⁷ are independently selected from hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkyl; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkylsulfonyl; cyanoC₁₋₆alkyl; hydroxyC₁₋₆alkyloxy; hydroxyC₁₋₆alkylamino; aminoC₁₋₆alkyloxy; di(C₁₋₆alkyl)aminocarbonyl; di(hydroxyC₁₋₆alkyl)amino; arylC₁₋₆alkyl)amino; di(C₁₋₆alkyl)aminoC₁₋₆alkyloxy; di(C₁₋₆alkyl)aminoC₁₋₆alkyloxy; arylSulfonylamino; aryloxy; arylC₂₋₆alkenediyl; di(C₁₋₆alkyl)amino; di(C₁₋₆alkyl)aminoC₁₋₆alkyl;

di(C₁₋₆alkyl)aminoC₁₋₆alkyl(C₁₋₆alkyl)aminoC₁₋₆alkyl; cyano; thiophenyl; thiophenyl substituted with di(C₁₋₆alkyl)aminoC₁₋₆alkyl(C₁₋₆alkyl)aminoC₁₋₆alkyl, di(C₁₋₆alkyl)aminoC₁₋₆alkyl, C₁₋₆alkylpiperazinylC₁₋₆alkyl or di(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkyl; furanyl; imidazolyl; C₁₋₆alkyltriazolyl; tetrazolyl; pyrrolidinyl; piperidinylC₁₋₆alkyloxy; morpholinyl; C₁₋₆alkylmorpholinyl; morpholinylC₁₋₆alkyloxy; morpholinylC₁₋₆alkylpiperazinyl; C₁₋₆alkylpiperazinylC₁₋₆alkyloxy; C₁₋₆alkylpiperazinylC₁₋₆alkyl; C₁₋₆alkylpiperazinylsulfonyl; aminosulfonylpiperazinylC₁₋₆alkyl; di(C₁₋₆alkyl)aminosulfonylpiperazinyl; di(C₁₋₆alkyl)aminosulfonylpiperazinyl; hydroxyC₁₋₆alkylpiperazinylC₁₋₆alkyl; C₁₋₆alkylpiperazinyl; hydroxyC₁₋₆alkylpiperazinylC₁₋₆alkyl; hydroxyC₁₋₆alkylpiperazinyl; hydroxyC₁₋₆alkylpiperazinylC₁₋₆alkylpiperazinylC₁₋₆alkylpiperazinyl; hydroxyC₁₋₆alkylpiperazinylC₁

pyrrolidinylC₁₋₆alkyloxy; pyrazolyl; thiopyrazolyl; pyrazolyl substituted with two substituents selected from C_{1.6}alkyl or trihaloC_{1.6}alkyl; pyridinyl; pyridinyl substituted with C₁₋₆alkyloxy or aryl; pyrimidinyl; quinolinyl; indole; phenyl; phenyl substituted with one, two or three substituents independently selected from halo, amino, C₁₋₆alkyl, C₁₋₆alkyloxy, hydroxyC₁₋₄alkyl, trifluoromethyl, trifluoromethyloxy, hydroxyC₁₋₄alkyloxy, C₁₋₄alkyloxyC₁₋₄alkyloxy, aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)amino, di(C_{1.4}alkyl)aminoC_{1.4}alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl(C₁₋₄alkyl)aminoC₁₋₄alkyl, piperidinylC₁₋₄alkyloxy, pyrrolidinylC_{1.4}alkyloxy, aminosulfonylpiperazinyl, aminosulfonylpiperazinylC₁₋₄alkyl, di(C₁₋₄alkyl)aminosulfonylpiperazinyl, di(C₁₋₄alkyl)aminosulfonylpiperazinylC₁₋₄alkyl, hydroxyC₁₋₄alkylpiperazinyl, hydroxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, C₁₋₄alkyloxypiperidinyl, C₁₋₄alkyloxypiperidinylC₁₋₄alkyl, hydroxyC₁₋₄alkyloxyC₁₋₄alkylpiperazinyl, hydroxyC₁₋₄alkyloxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, (hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)amino, (hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)aminoC₁₋₄alkyl, pyrrolidinylC₁₋₄alkyloxy, morpholinylC₁₋₄alkyloxy, morpholinylC₁₋₄alkyl, C₁₋₄alkylpiperazinyl, C₁₋₄alkylpiperazinylC₁₋₄alkyloxy, C₁₋₄alkylpiperazinylC₁₋₄alkyl, hydroxyC₁₋₄alkylamino, di(hydroxyC₁₋₄alkyl)amino, di(C₁₋₄alkyl)aminoC₁₋₄alkylamino, aminothiadiazolyl, aminosulfonylpiperazinylC₁₋₄alkyloxy, or thiophenylC₁₋₄alkylamino.

146. The compound of claim 143 wherein:

t = 0;

m = 0:

is hydrogen, halo, hydroxy, amino, nitro, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl or di(C₁₋₆alkyl)amino;

- -L- is a direct bond or a bivalent radical selected from C_{1-6} alkanediyl, C_{1-6} alkanediyloxy, amino or carbonyl;
- R¹⁴ is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyloxy, arylC₁₋₆alkyl, aminocarbonyl, aminoC₁₋₆alkyl, C₁₋₆alkylaminoC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl;
- R¹⁵ is hydrogen;
 - is a radical selected from (a-1), (a-3), (a-4), (a-5), (a-6), (a-7), (a-8), (a-9), (a-10), (a-11), (a-12), (a-13), (a-14), (a-15), (a-16), (a-17), (a-18), (a-19), (a-20), (a-21), (a-22), (a-23), (a-24), (a-25), (a-26), (a-28), (a-29), (a-30), (a-31), (a-32), (a-33), (a-34), (a-35), (a-36), (a-37), (a-38), (a-39), (a-40), (a-41), (a-42), (a-44), (a-45), (a-46), (a-47), (a-48) or (a-51); each s is independently 0, 1, 2, 3 or 4;
- R⁶ is hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkyloxy; di(C₁₋₆alkyl)amino; cyano; thiophenyl; furanyl; furanyl substituted with hydroxyC₁₋₆alkyl; benzofuranyl; imidazolyl; oxazolyl; oxazolyl substituted with aryl and C₁₋₆alkyl; C₁₋₆alkyltriazolyl; tetrazolyl; pyrrolidinyl; pyrrolyl; morpholinyl; C₁₋₆alkylmorpholinyl; piperazinyl; C₁₋₆alkylpiperazinyl; C₁₋₆alkylpiperazinyl; C₁₋₆alkylpiperazinyl; pyrazolyl substituted with one or two substituents selected from C₁₋₆alkyl or trihaloC₁₋₆alkyl; pyridinyl; pyridinyl substituted with C₁₋₆alkyloxy, aryloxy or aryl; pyrimidinyl; quinolinyl; indole; phenyl; or phenyl substituted with one or two substituents independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy or trifluoromethyl;

 R^7 is hydrogen; halo; hydroxy; amino; nitro; trihalo C_{1-6} alkyl; trihalo C_{1-6} alkyloxy; C_{1-6} alkyl; C_{1-6} alkyloxy; C_{1-6} alkylcarbonyl; C_{1-6} alkyloxycarbonyl; C_{1-6} alkylsulfonyl; hydroxy C_{1-6} alkyl; aryloxy; di(C_{1-6} alkyl)amino; cyano; pyridinyl; phenyl; or phenyl substituted with one or two substituents independently selected from halo, C_{1-6} alkyl, C_{1-6} alkyloxy or trifluoromethyl.

147. The compound of claim 143 wherein:

n is 1; m is 0 or 1; t is 0, 1 or 2; Q is —C, or —CH, or —CH, or —CH, ; R¹² is hydrogen; -L- is a direct bond; R¹⁴ and R¹⁵ are H;

is a radical selected from (a-1), (a-20), (a-27), (a-28), (a-29), (a-41) or (a-42); each s is independently 0, 1 or 2; and each R⁶ is independently selected from hydrogen, halo, C₁₋₆alkyl or C₁₋₆alkyloxy.

148. The compound of claim 143 that is selected from one of

wherein the terminal hydroxamic acid moiety (-C(O)-NH-OH) is replaced with

wherein Φ , R¹, R², R³, and R⁴ are as defined in accordance with claim 1.

- 149. The compound of claim 143 wherein R¹, R², R³, and R⁴ are all H.
- 150. A compound according to claim 143 for use in inhibting histone deacetylase.
- 151. A compound according to calim 143 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.

152. The compound of claim 151, wherein said treatment is effected by inhibiting histone deacetylase.

- 153. The compound of calim 151, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 154. The compound of claim 151, wherein said cell proliferative disease is cancer.
- 155. The compound of claim 154, wherein said cancer is a solid tumor cancer.
- 156. The compound of claim 154, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 157. A pharmaceutical composition comprising a compound according to claim 143 and a pharmaceutically acceptable carrier.
- 158. The pharmaceutical composition of claim 157 further comprising a nucleic acid level inhibitor of histone deacetylase.
- 159. The pharmaceutical composition of claim 158, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 160. The pharmaceutical composition of claim 159, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:I, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:IO, SEQ ID No:II, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.
- 161. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 143.
- 162. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 157
- 163. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 158.

164. The method of claim 162, wherein said cell proliferative disease is a neoplastic cell proliferative disease.

- 165. The method of claim 162, wherein said cell proliferative disease is cancer.
- 166. The method of claim 165, wherein said cancer is a solid tumor cancer.
- 167. The method of claim 166, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 168. The method of claim 163, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 169. The method of claim 163, wherein said cell proliferative disease is cancer.
- 170. The method of claim 169, wherein said cancer is a solid tumor cancer.
- 171. The method of claim 170, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 172. A compound of the formula:

$$R^{1}$$
 R^{2}
 $R^{3} = A$
 R^{4}
 R^{4}
 R^{4}
 R^{14}
 $R^{$

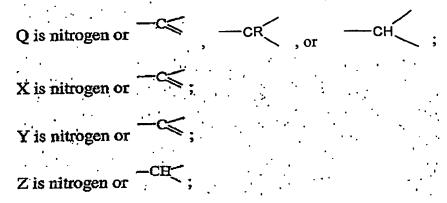
or a pharmaceutically acceptable salt thereof, wherein

 Φ is -NH $_2$ or -OH;

R1 is H or as defined in claim 1;

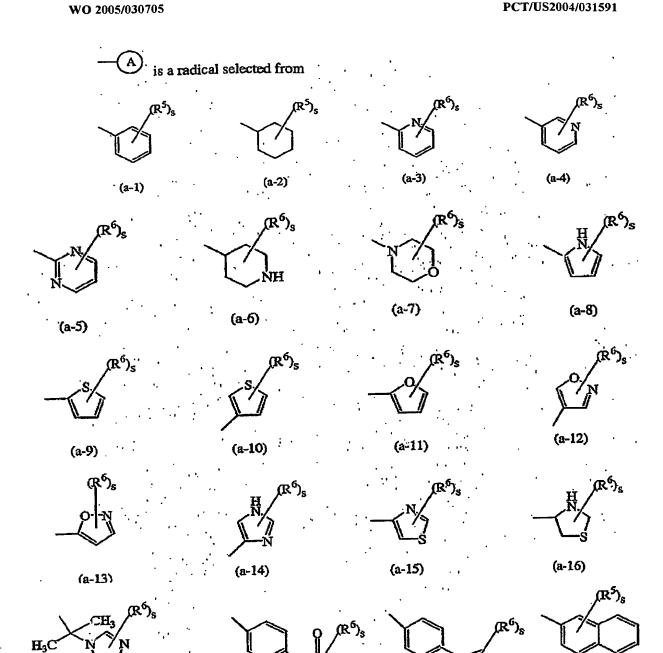
R2, R3, and R4 are as defined in claim 1;

n is 0, 1, 2 or 3 and when n is 0 then a direct bond is intended; t is 0, 1, 2, 3 or 4 and when t is 0 then a direct bond is intended;



R is selected from the group consisting of hydrogen, halogen, -NH₂, nitro, hydroxy, aryl, heterocyclyl, C_3 - C_8 -cycloalkyl, heteroaryl, C_1 - C_7 -akyl, haloalkyl, C_1 - C_7 -alkenyl, C_1 - C_7 -alkyl-aryloxy, C_1 - C_7 -alkyl-arylsulfanyl, C_1 - C_7 -alkyl-arylsulfinyl, C_1 - C_7 -alkyl-arylaminosulfonyl, C_1 - C_7 -alkyl-arylamine, C_1 - C_7 -alkyl or C_1 - C_7 -alkyl-arylamine, C_1 - C_7 -alkyl or C_1 - C_7 -alkoxy;

- R¹² is hydrogen, halo, hydroxy, amino, nitro, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl, di(C₁₋₆alkyl)amino, hydroxyamino or naphtalenylsulfonylpyrazinyl;
- -L- is a direct bond or a bivalent radical selected from C₁₋₆alkanediyl, C₁₋₆alkyloxy, amino, carbonyl or aminocarbonyl;
- each R¹³ independently represents a hydrogen atom and one hydrogen atom can be replaced by a substituent selected from aryl;
- R¹⁴ is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyloxy, arylC₁₋₆alkyl, aminocarbonyl, hydroxycarbonyl, aminoC₁₋₆alkyl, aminocarbonylC₁₋₆alkyl, hydroxycarbonylC₁₋₆alkyl, hydroxyaminocarbonyl, C₁₋₆alkyloxycarbonyl, C₁₋₆alkylaminoC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl;

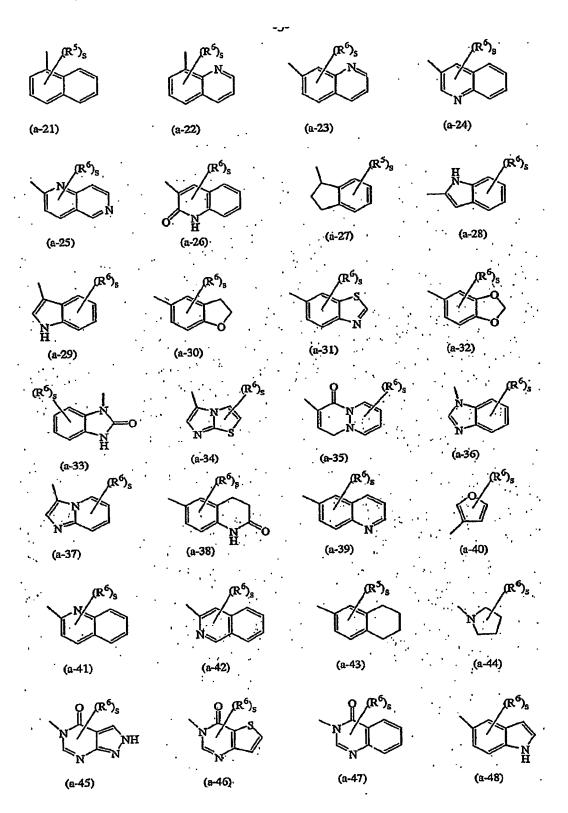


(a-18)

(a-17)

(a-19)

(a-20)



$$(a-49)$$
 $(a-50)$ $(R^6)_s$ $(R^6)_s$ $(R^6)_s$ $(a-51)$

wherein each s is independently 0, 1, 2, 3, 4 or 5;

are independently selected from hydrogen; halo; hydroxy; amino; nitro; each R5 and R6 trihaloC1-6alkyl; trihaloC1-6alkyloxy; C1-6alkyl; C1-6alkyl substituted with aryl and $C_{3\text{-}10} \text{cycloalkyl}; \ C_{1\text{-}6} \text{alkyloxy}; \ C_{1\text{-}6} \text{alkyloxy} C_{1\text{-}6} \text{alkyloxy}; \ C_{1\text{-}6} \text{alkyloxy}; C_{1\text{-}6} \text{alkylox$ C_{1-6} alkyloxycarbonyl; C_{1-6} alkylsulfonyl; cyano C_{1-6} alkyl; hydroxy C_{1-6} alkyl; $\label{eq:hydroxyC1-6} \text{hydroxyC}_{1\text{-}6} \text{alkyloxy; hydroxyC}_{1\text{-}6} \text{alkyloxy; aminoC}_{1\text{-}6} \text{alkyloxy;} \\$ $\label{eq:condition} \begin{array}{l} di(C_{1\text{-}6}alkyl) amino; (aryl)(C_{1\text{-}6}alkyl) amino; (aryl)(C_{1\text{-}6}alkyl) amino; \\ \end{array}$ di(C₁₋₆alkyl)aminoC₁₋₆alkyloxy; di(C₁₋₆alkyl)aminoC₁₋₆alkylamino; $di(C_{1-6}alkyl)aminoC_{1-6}alkylaminoC_{1-6}alkyl;$ arylsulfonyl; arylsulfonylamino; aryloxy; aryloxyC₁₋₆alkyl; arylC₂₋₆alkenediyl; di(C₁₋₆alkyl)amino; di(C₁₋₆alkyl)aminoC₁₋₆alkyl; di(C₁₋₆alkyl)amino(C₁₋₆alkyl)amino; di(C_{1.6}alkyl)amino(C_{1.6}alkyl)aminoC_{1.6}alkyl; $di(C_{1-6}alkyl)aminoC_{1-6}alkyl(C_{1-6}alkyl)amino;$ $di(C_{1-6}alkyl)aminoC_{1-6}alkyl(C_{1-6}alkyl)aminoC_{1-6}alkyl;$ aminosulfonylamino(C₁₋₆alkyl)amino; aminosulfonylamino(C₁₋₆alkyl)aminoC₁₋₆alkyl; di(C₁₋₆alkyl)aminosulfonylamino(C₁₋₆alkyl)amino;

di(C₁₋₆alkyl)aminosulfonylamino(C₁₋₆alkyl)aminoC₁₋₆alkyl; cyano; thiophenyl; thiophenyl substituted with $di(C_{1-6}alkyl)aminoC_{1-6}alkyl(C_{1-6}alkyl)aminoC_{1-6}alkyl,$ $di(C_{1-6}alkyl)aminoC_{1-6}alkyl, C_{1-6}alkylpiperazinylC_{1-6}alkyl,$ hydroxyC₁₋₆alkylpiperazinylC₁₋₆alkyl, hydroxyC₁₋₆alkyloxyC₁₋₆alkylpiperazinylC₁₋₆alkyl, di(C₁₋₆alkyl)aminosulfonylpiperazinylC₁₋₆alkyl, C1-6alkyloxypiperidinyl, C1-6alkyloxypiperidinylC1-6alkyl, morpholinylC1-6alkyl, hydroxyC_{1.6}alkyl(C_{1.6}alkyl)aminoC_{1.6}alkyl, or di(hydroxyC_{1.6}alkyl)aminoC_{1.6}alkyl; furanyl; furanyl substituted with hydroxyC₁₋₆alkyl; benzofuranyl; imidazolyl; oxazolyl; oxazolyl substituted with aryl and C₁₋₆alkyl; C₁₋₆alkyltriazolyl; tetrazolyl; pyrrolidinyl; pyrrolyl; piperidinylC₁₋₆alkyloxy; morpholinyl; C₁₋₆alkylmorpholinyl; morpholinylC₁₋₆alkyloxy; morpholinylC₁₋₆alkyl; morpholinylC₁₋₆alkylamino; morpholinylC₁₋₆alkylaminoC₁₋₆alkyl; piperazinyl; C₁₋₆alkylpiperazinyl; C₁₋₆alkylpiperazinylC₁₋₆alkyloxy; piperazinylC₁₋₆alkyl; naphtalenyisulfonylpiperazinyl; naphtalenylsulfonylpiperidinyl; naphtalenylsulfonyl; C_{1-6} alkylpiperazinyl C_{1-6} alkyl; C_{1-6} alkylpiperazinyl C_{1-6} alkylamino; C_{1-6} alkylpiperazinyl C_{1-6} alkylamino C_{1-6} alkyl; C_{1-6} alkylpiperazinylsulfonyl; aminosulfonylpiperazinylC₁₋₆alkyloxy; aminosulfonylpiperazinyl; $aminosulfonylpiperazinyl C_{1-6} alkyl; \ di(C_{1-6} alkyl) aminosulfonylpiperazinyl;$ $\label{eq:condition} di(C_{1\text{-}6}alkyl) aminosulfonyl piperazinyl C_{1\text{-}6}alkyl; \ hydroxyC_{1\text{-}6}alkyl piperazinyl;$ $hydroxyC_{1-6}alkylpiperazinylC_{1-6}alkyl; C_{1-6}alkyloxypiperidinyl;$ $C_{1\text{--}6}$ alkyloxypiperidinyl $C_{1\text{--}6}$ alkyl; piperidinylamino $C_{1\text{--}6}$ alkylamino; piperidinylaminoC₁₋₆alkylaminoC₁₋₆alkyl; (C₁₋₆alkylpiperidinyl)(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkylamino; $(C_{1-6}alkylpiperidinyl)(hydroxyC_{1-6}alkyl)aminoC_{1-6}alkylaminoC_{1-6}alkyl;$ hydroxyC₁₋₆alkyloxyC₁₋₆alkylpiperazinyl; $hydroxyC_{1-6}alkyloxyC_{1-6}alkylpiperazinylC_{1-6}alkyl;\\$ $(hydroxyC_{1-6}alkyl)(C_{1-6}alkyl)amino; (hydroxyC_{1-6}alkyl)(C_{1-6}alkyl)aminoC_{1-6}alkyl;$

hydroxyC₁₋₆alkylaminoC₁₋₆alkyl; di(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkyl; pyrrolidinylC₁₋₆alkyl; pyrrolidinylC₁₋₆alkyloxy; pyrazolyl; thiopyrazolyl; pyrazolyl substituted with two substituents selected from C1-6alkyl or trihaloC1-6alkyl; pyridinyl; pyridinyl substituted with C₁₋₆alkyloxy, aryloxy or aryl; pyrimidinyl; tetrahydropyrimidinylpiperazinyl; tetrahydropyrimidinylpiperazinylC1_6alkyl; quinolinyl; indole; phenyl; phenyl substituted with one, two or three substituents independently selected from halo, amino, nitro, C1-6alkyl, C1-6alkyloxy, hydroxyC₁₋₄alkyl, trifluoromethyl, trifluoromethyloxy, hydroxyC₁₋₄alkyloxy, C₁₋₄alkylsulfonyl, C₁₋₄alkyloxyC₁₋₄alkyloxy, C₁₋₄alkyloxycarbonyl, aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)aminocarbonyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)amino(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl(C₁₋₄alkyl)amino, $di(C_{1-4}alkyl)aminoC_{1-4}alkyl(C_{1-4}alkyl)aminoC_{1-4}alkyl,$ aminosulfonylamino(C1-4alkyl)amino, aminosulfonylamino(C1-4alkyl)aminoC1-4alkyl, di(C1-4alkyl)aminosulfonylamino(C1-4alkyl)amino, di(C1_4alkyl)aminosulfonylamino(C1_4alkyl)aminoC1_6alkyl, cyano, piperidinylC₁₋₄alkyloxy, pyrrolidinylC₁₋₄alkyloxy, aminosulfonylpiperazinyl, aminosulfonylpiperazinylC₁₋₄alkyl, di(C₁₋₄alkyl)aminosulfonylpiperazinyl, di(C₁₋₄alkyl)aminosulfonylpiperazinylC₁₋₄alkyl, hydroxyC₁₋₄alkylpiperazinyl, hydroxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, C₁₋₄alkyloxypiperidinyl,

C₁₋₄alkyloxypiperidinylC₁₋₄alkyl, hydroxyC₁₋₄alkyloxyC₁₋₄alkylpiperazinyl, hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)amino, (hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)aminoC₁₋₄alkyl, (hydroxyC₁₋₄alkyl)aminoC₁₋₄alkyl, di(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkyl, furanyl, furanyl substituted with -CH=CH-CH=CH-, pyrrolidinylC₁₋₄alkyl, pyrrolidinylC₁₋₄alkyloxy, morpholinyl, morpholinylC₁₋₄alkyloxy, morpholinylC₁₋₄alkylamino, morpholinylC₁₋₄alkylaminoC₁₋₄alkyl, piperazinyl, C₁₋₄alkylpiperazinyl, C₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperazinylC₁₋₄alkylpiperidinyl)(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkylamino, (C₁₋₄alkylpiperidinyl)(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkylaminoC₁₋₄alkyl, pyridinylC₁₋₄alkyloxy,

hydroxyC₁₋₄alkylamino, hydroxyC₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkylamino, aminothiadiazolyl, aminosulfonylpiperazinylC₁₋₄alkyloxy, or thiophenylC₁₋₄alkylamino;

each R5 and R6 can be placed on the nitrogen in replacement of the hydrogen;

aryl in the above is phenyl, or phenyl substituted with one or more substituents each independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl, cyano or hydroxycarbonyl.

173. The compound of claim 172 wherein each of R², R³, and R⁴ corresponds to R¹², R¹³, and R¹⁴, respectively, in claim 172 wherein:

n is 1 or 2; t is 0, 1, 2 or 4; Q is

R² is hydrogen or nitro;

-L- is a direct bond or a bivalent radical selected from C₁₋₆alkanediyl;

R⁴ is hydrogen;

(a-20), (a-32), (a-33), (a-47) or (a-51);

each s is independently 0, 1, 2, or 4;

each R⁵ and R⁶ are independently selected from hydrogen; halo; trihaloC₁₋₆alkyl; C₁₋₆alkyl; C₁₋₆alkyl; C₁₋₆alkyl; C₁₋₆alkyloxy; C₁₋₆alkylcarbonyl; benzofuranyl; naphtalenylsulfonyl; pyridinyl substituted with aryloxy; phenyl; or phenyl substituted with one substituent independently selected from hydroxyC₁₋₄alkyl or morpholinylC₁₋₄alkyl.

174. The compound of claim 170 wherein each of R², R³, and R⁴ corresponds to R¹², R¹³, and R¹⁴, respectively, in claim 172 wherein:

n is 1;

R² is hydrogen;

-L- is a direct bond;

each R3 independently represents a hydrogen atom;

R⁴ is hydrogen;

is a radical selected from (a-6), (a-11), (a-20), (a-47) or (a-51); each s is independently 0, 1, or 4;

each R⁵ and R⁶ are independently selected from hydrogen; C₁₋₆alkyl; C₁₋₆alkyloxy; naphtalenylsulfonyl; or phenyl substituted with hydroxyC₁₋₄alkyl or morpholinylC₁₋₄alkyl.

175. The compound of claim 172 wherein L is a direct bond.

176. The compound of claim 172 wherein each of R^2 , R^3 , and R^4 corresponds to R^{12} , R^{13} , and R^{14} , respectively, in claim 172 wherein : t is 1, 2, 3, or 4;

- R² is hydrogen, halo, hydroxy, amino, nitro, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl or di(C₁₋₆alkyl)amino;
- -L- is a direct bond or a bivalent radical selected from C_{1-6} alkanediyl, C_{1-6} alkanediyloxy, amino or carbonyl;
- R⁴ is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyl, aminocarbonyl, aminoC₁₋₆alkyl, C₁₋₆alkylaminoC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl;
- (a-10), (a-11), (a-12), (a-13), (a-14), (a-15), (a-16), (a-17), (a-18), (a-19), (a-10), (a-11), (a-12), (a-13), (a-14), (a-15), (a-16), (a-17), (a-18), (a-19), (a-20), (a-21), (a-22), (a-23), (a-24), (a-25), (a-26), (a-28), (a-29), (a-30), (a-31), (a-32), (a-33), (a-34), (a-35), (a-36), (a-37), (a-38), (a-39), (a-40), (a-41), (a-42), (a-44), (a-45), (a-46), (a-47), (a-48) and (a-51); each s is independently 0, 1, 2, 3 or 4;

each s is independently 0, 1, 2, 3 or 4;

R⁵ is hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkyl; C₁₋₆alkyloxy; C₁₋₆alkylcarbonyl; C₁₋₆alkyloxycarbonyl; C₁₋₆alkylsulfonyl; hydroxyC₁₋₆alkyl; aryloxy; di(C₁₋₆alkyl)amino; cyano; thiophenyl; furanyl; furanyl substituted with hydroxyC₁₋₆alkyl; benzofuranyl; imidazolyl; oxazolyl; oxazolyl substituted with aryl and C₁₋₆alkyl; C₁₋₆alkyltriazolyl; tetrazolyl; pyrrolidinyl; pyrrolyl; morpholinyl; C₁₋₆alkylmorpholinyl; piperazinyl;

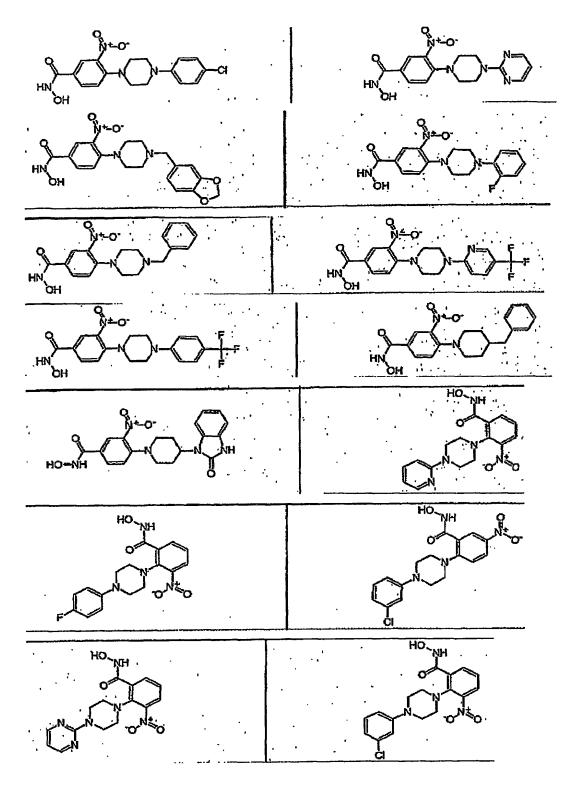
C₁₋₆alkylpiperazinyl; hydroxyC₁₋₆alkylpiperazinyl;

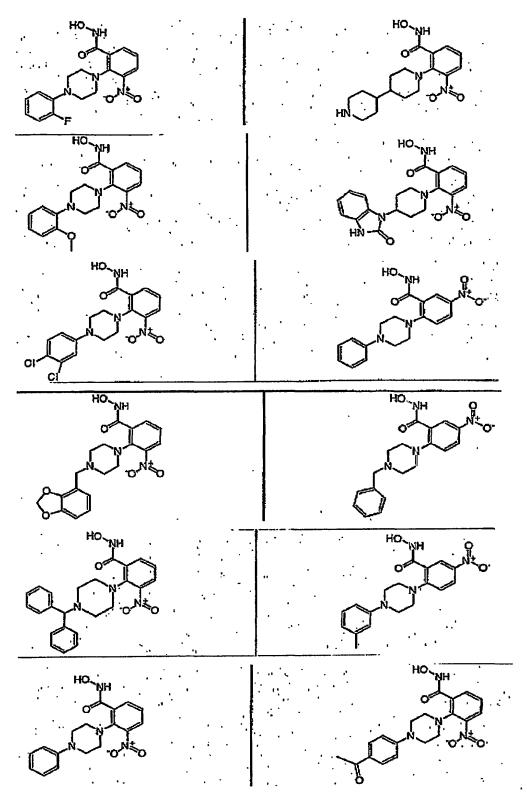
 C_{1-6} alkyloxypiperidinyl; pyrazoly; pyrazolyl substituted with one or two substituents selected from C_{1-6} alkyl or trihalo C_{1-6} alkyl; pyridinyl; pyridinyl substituted with C_{1-6} alkyloxy, aryloxy or aryl; pyrimidinyl; quinolinyl; indole; phenyl; or phenyl substituted with one or two substituents independently selected from halo, C_{1-6} alkyl, C_{1-6} alkyloxy or trifluoromethyl;

 R^6 is hydrogen; halo; hydroxy; amino; nitro; trihalo C_{1-6} alkyl; trihalo C_{1-6} alkyloxy; C_{1-6} alkyloxy; C_{1-6} alkyloxy; C_{1-6} alkyloxycarbonyl; C_{1-6} alkyloxycarbonyl;

 C_{1-6} alkylsulfonyl; hydroxy C_{1-6} alkyl; aryloxy; di(C_{1-6} alkyl)amino; cyano; pyridinyl; phenyl; or phenyl substituted with one or two substituents independently selected from halo, C_{1-6} alkyl, C_{1-6} alkyloxy or trifluoromethyl.

177. The compound of claim 172 that is selected from one of





wherein the terminal hydroxamic acid moiety (-C(O)-NH-OH) is replaced with

wherein Φ , R^1 , R^2 , R^3 , and R^4 are as defined in accordance with claim 1.

178. The compound of claim 172 wherein R1, R2, R3, and R4 are all H.

179. A compound according to claim 172 for use in inhibting histone deacetylase.

180. A compound according to calim 172 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.

181. The compound of claim 180, wherein said treatment is effected by inhibiting histone deacetylase.

182. The compound of calim 180, wherein said cell proliferative disease is a neoplastic cell proliferative disease.

183. The compound of claim 180, wherein said cell proliferative disease is cancer.

184. The compound of claim 183, wherein said cancer is a solid tumor cancer.

185. The compound of claim 183, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.

- 186. A pharmaceutical composition comprising a compound according to claim 172 and a pharmaceutically acceptable carrier.
- 187. The pharmaceutical composition of claim 186 further comprising a nucleic acid level inhibitor of histone deacetylase.
- 188. The pharmaceutical composition of claim 187, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 189. The pharmaceutical composition of claim 188, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:I, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:II, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.
- 190. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 172.
- 191. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 186.
- 192. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 187.
- 193. The method of claim 191, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 194. The method of claim 191, wherein said cell proliferative disease is cancer.
- 195. The method of claim 194, wherein said cancer is a solid tumor cancer.
- 196. The method of claim 195, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.

197. The method of claim 192, wherein said cell proliferative disease is a neoplastic cell proliferative disease.

- 198. The method of claim 192, wherein said cell proliferative disease is cancer.
- 199. The method of claim 198, wherein said cancer is a solid tumor cancer.
- 200. The method of claim 199, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 201. A compound of the formula:

or a pharmaceutically acceptable salt thereof, wherein

 Φ is -NH 2 or -OH;

R1 is H or as defined in claim 1;

R2, R3, and R4 are as defined in claim 1;

in is 0, I, 2 or 3 and when n is Qthen a direct bond is intended;

t is 0, 1, 2, 3 or 4 and when t is 0 then a direct bond is intended;

Q is nitrogen or
$$-CR$$
, $-CR$, α

X is nitrogen or —C

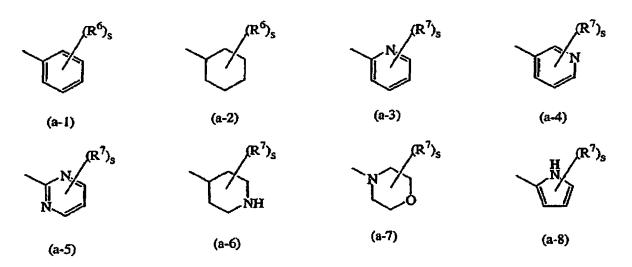
Y is nitrogen or —

Z is nitrogen or —CH*"*

R is selected from the group consisting of hydrogen, halogen, -NH₂, nitro, hydroxy, aryl, heterocyclyl, $C_3 \cdot C_8$ -cycloalkyl, heteroaryl, $C_r \cdot C_7$ -akyl, haloalkyl, CrC_7 -alkenyl, $C_r \cdot C_7$ -alkyl-arylsulfanyl, $C_r \cdot C_7$ -alkyl-arylsulfanyl, $C_r \cdot C_7$ -alkyl-arylsulfonyl, $C_r \cdot C_7$ -alkyl-arylsulfonyl, C

arylaminosulfonyl, C₁-C₇-alkyl-arylamine, C₁-C₇-alkynyl-C(0)-amine, C₁-C₇-alkenyl-C(0)-amine, C₁-C₇-alkynyl-R⁹, C₁-C₇-alkenyl-R⁹ wherein R⁹ is hydrogen , hydroxy, amino, C₁-C₇-alkyl or C₁-C₇-alkoxy;

- each R¹² 1ydrogen, halo, hydroxy, amino, nitro, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl, di(C₁₋₆alkyl)amino, hydroxyamino or naphtalenylsulfonylpyrazinyl;
 - each R¹³ independently represents a hydrogen atom and one hydrogen atom can be replaced by a substituent selected from aryl;
 - R¹⁴ is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyloxy, arylC₁₋₆alkyl, aminocarbonyl, hydroxycarbonyl, aminoC₁₋₆alkyl, aminocarbonylC₁₋₆alkyl, hydroxycarbonylC₁₋₆alkyl, hydroxyaminocarbonyl, C₁₋₆alkyloxycarbonyl, C₁₋₆alkylaminoC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl;
 - R¹⁵ is hydrogen, C₁₋₆alkyl, C₃₋₁₀cycloalkyl, hydroxyC₁₋₆alkyl, C₁₋₆alkyloxyC₁₋₆alkyl, di(C₁₋₆alkyl)aminoC₁₋₆alkyl or aryl;
 - is a radical selected from



$$(a-9) \qquad (a-10) \qquad (a-11) \qquad (a-12)$$

$$(a-13) \qquad (a-14) \qquad (a-15) \qquad (a-16)$$

$$(a-17) \qquad (a-18) \qquad N \qquad (a-19) \qquad N \qquad (a-20)$$

$$(a-21) \qquad (a-22) \qquad (a-23) \qquad (a-24)$$

$$(a-25) \qquad (a-26) \qquad (a-27) \qquad (a-28)$$

$$(a-49)$$
 $(a-50)$ $(R^7)_S$ $(R^7)_S$ $(R^7)_S$ $(a-51)$

wherein each s is independently 0, 1, 2, 3, 4 or 5;

each R⁶ and R⁷ are independently selected from hydrogen; halo; hydroxy; amino; nitro;

 $trihaloC_{1-6}$ alkyl; $trihaloC_{1-6}$ alkyloxy; C_{1-6} alkyl; C_{1-6} alkyl substituted with aryl and

C₃₋₁₀cycloalkyl; C₁₋₆alkyloxy; C₁₋₆alkyloxyC₁₋₆alkyloxy; C₁₋₆alkylcarbonyl;

C₁₋₆alkyloxycarbonyl; C₁₋₆alkylsulfonyl; cyanoC₁₋₆alkyl; hydroxyC₁₋₆alkyl;

hydroxyC₁₋₆alkyloxy; hydroxyC₁₋₆alkylamino; aminoC₁₋₆alkyloxy;

di(C₁₋₆alkyl)aminocarbonyl; di(hydroxyC₁₋₆alkyl)amino; (aryl)(C₁₋₆alkyl)amino;

di(C₁₋₆alkyl)aminoC₁₋₆alkyloxy; di(C₁₋₆alkyl)aminoC₁₋₆alkylamino;

 $di(C_{1\text{-}6}alkyl)aminoC_{1\text{-}6}alkylaminoC_{1\text{-}6}alkyl;\ arylsulfonylamino;$

aryloxy; aryloxyC₁₋₆alkyl; arylC₂₋₆alkenediyl; di(C₁₋₆alkyl)amino;

 $di(C_{1\text{-}6}alkyl)aminoC_{1\text{-}6}alkyl; di(C_{1\text{-}6}alkyl)amino(C_{1\text{-}6}alkyl)amino;\\$

di(C₁₋₆alkyl)amino(C₁₋₆alkyl)aminoC₁₋₆alkyl;

 $di(C_{1-6}alkyl)aminoC_{1-6}alkyl(C_{1-6}alkyl)amino;$

 $di(C_{1-6}alkyl)aminoC_{1-6}alkyl(C_{1-6}alkyl)aminoC_{1-6}alkyl;$

aminosulfonylamino(C1-6alkyl)amino;

arninosulfonylamino(C₁₋₆alkyl)aminoC₁₋₆alkyl;

di(C₁₋₆alkyl)aminosulfonylamino(C₁₋₆alkyl)amino;

 $di(C_{1-6}alkyl)$ aminosulfonylamino $(C_{1-6}alkyl)$ amino $C_{1-6}alkyl$; cyano; thiophenyl;

thiophenyl substituted with $di(C_{1-6}alkyl)aminoC_{1-6}alkyl(C_{1-6}alkyl)aminoC_{1-6}alkyl,$ $di(C_{1-6}alkyl)aminoC_{1-6}alkyl,$ $C_{1-6}alkylpiperazinylC_{1-6}alkyl,$

hydroxyC₁₋₆alkylpiperazinylC₁₋₆alkyl,
hydroxyC₁₋₆alkylpiperazinylC₁₋₆alkyl,
di(C₁₋₆alkyl)aminosulfonylpiperazinylC₁₋₆alkyl,
C₁₋₆alkyloxypiperidinyl, C₁₋₆alkyloxypiperidinylC₁₋₆alkyl, morpholinylC₁₋₆alkyl,
hydroxyC₁₋₆alkyl(C₁₋₆alkyl)aminoC₁₋₆alkyl, or di(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkyl;
furanyl; furanyl substituted with hydroxyC₁₋₆alkyl; benzofuranyl; imidazolyl;
oxazolyl; oxazolyl substituted with aryl and C₁₋₆alkyl; C₁₋₆alkyltriazolyl; tetrazolyl;
pyrrolidinyl; pyrrolyl; piperidinylC₁₋₆alkyloxy; morpholinyl; C₁₋₆alkylmorpholinyl;
morpholinylC₁₋₆alkyloxy; morpholinylC₁₋₆alkyl; morpholinylC₁₋₆alkylamino;
morpholinylC₁₋₆alkylaminoC₁₋₆alkyl; piperazinyl; C₁₋₆alkylpiperazinyl;
C₁₋₆alkylpiperazinylC₁₋₆alkyloxy; piperazinylC₁₋₆alkyl;
naphtalenylsulfonylpiperazinyl; naphtalenylsulfonylpiperidinyl; naphtalenylsulfonyl;

C_{1.6}alkylpiperazinylC_{1.6}alkyl; C_{1.6}alkylpiperazinylC_{1.6}alkylamino; C_{1.6}alkylpiperazinylC_{1.6}alkylaminoC_{1.6}alkyl; C_{1.6}alkylpiperazinylsulfonyl; aminosulfonylpiperazinylC_{1.6}alkyloxy; aminosulfonylpiperazinyl; aminosulfonylpiperazinylC₁₋₆alkyl; di(C₁₋₆alkyl)aminosulfonylpiperazinyl; di(C_{1.6}alkyl)aminosulfonylpiperazinylC_{1.6}alkyl; hydroxyC_{1.6}alkylpiperazinyl; hydroxyC₁₋₆alkylpiperazinylC₁₋₆alkyl; C₁₋₆alkyloxypiperidinyl; C_{1-6} alkyloxypiperidinyl C_{1-6} alkyl; piperidinylamino C_{1-6} alkylamino; piperidinylaminoC₁₋₆alkylaminoC₁₋₆alkyl; (C₁₋₆alkylpiperidinyl)(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkylamino; (C₁₋₆alkylpiperidinyl)(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkylaminoC₁₋₆alkyl; hydroxyC_{1.6}alkyloxyC_{1.6}alkylpiperazinyl; hydroxyC₁₋₆alkyloxyC₁₋₆alkylpiperazinylC₁₋₆alkyl; $(hydroxyC_{1-6}alkyl)(C_{1-6}alkyl)amino; (hydroxyC_{1-6}alkyl)(C_{1-6}alkyl)aminoC_{1-6}alkyl;$ hydroxyC₁₋₆alkylaminoC₁₋₆alkyl; di(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkyl; pyrrolidinylC₁₋₆alkyl; pyrrolidinylC₁₋₆alkyloxy; pyrazolyl; thiopyrazolyl; pyrazolyl substituted with two substituents selected from C₁₋₆alkyl or trihaloC₁₋₆alkyl; pyridinyl; pyridinyl substituted with C_{1-6} alkyloxy, aryloxy or aryl; pyrimidinyl; tetrahydropyrimidinylpiperazinyl; tetrahydropyrimidinylpiperazinylC₁₋₆alkyl; quinolinyl; indole; phenyl; phenyl substituted with one, two or three substituents independently selected from halo, amino, nitro, C₁₋₆alkyl, C₁₋₆alkyloxy,

C₁₋₄alkylsulfonyl, C₁₋₄alkyloxyC₁₋₄alkyloxy, C₁₋₄alkyloxycarbonyl, aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)aminoC₁₋₄alkyl)aminoC₁₋₄alkyl)aminoC₁₋₄alkyl)aminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)

 $hydroxyC_{1-4}alkyl$, trifluoromethyl, trifluoromethyloxy, $hydroxyC_{1-4}alkyloxy$,

$$\label{eq:continuous_continuous_continuous} \begin{split} & \text{di}(C_{1\text{-4}}\text{alkyl})\text{aminosulfonylamino}(C_{1\text{-4}}\text{alkyl})\text{amino}C_{1\text{-6}}\text{alkyl}, \, \text{cyano}, \\ & \text{di}(C_{1\text{-4}}\text{alkyl})\text{aminosulfonylamino}(C_{1\text{-4}}\text{alkyl})\text{aminosulfonylpiperazinyl}, \\ & \text{piperidinyl}C_{1\text{-4}}\text{alkyloxy}, \, \text{pyrrolidinyl}C_{1\text{-4}}\text{alkyloxy}, \, \text{aminosulfonylpiperazinyl}, \\ & \text{aminosulfonylpiperazinyl}C_{1\text{-4}}\text{alkyl}, \, \text{di}(C_{1\text{-4}}\text{alkyl})\text{aminosulfonylpiperazinyl}, \\ & \text{di}(C_{1\text{-4}}\text{alkyl})\text{aminosulfonylpiperazinyl}C_{1\text{-4}}\text{alkyl}, \, \text{hydroxy}C_{1\text{-4}}\text{alkylpiperazinyl}, \\ & \text{hydroxy}C_{1\text{-4}}\text{alkylpiperazinyl}C_{1\text{-4}}\text{alkyl}, \, \text{hydroxy}C_{1\text{-4}}\text{alkylpiperazinyl}, \\ & \text{hydroxy}C_{1\text{-4}}\text{alkylpiperazinyl}C_{1\text{-4}}\text{alkyl}, \, \text{hydroxy}C_{1\text{-4}}\text{alkylpiperazinyl}, \\ & \text{hydroxy}C_{1\text{-4}}\text{alkylpiperazinyl}C_{1\text{-4}}\text{alkyl}, \, \\ \end{aligned}$$

(hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)amino, (hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(hydroxyC₁₋₄alkyl)amino, di(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkyl, furanyl, furanyl substituted with -CH=CH-CH=CH-, pyrrolidinylC₁₋₄alkyl, pyrrolidinylC₁₋₄alkyloxy, morpholinyl, morpholinylC₁₋₄alkyloxy, morpholinylC₁₋₄alkyl, morpholinylC₁₋₄alkylamino, morpholinylC₁₋₄alkylaminoC₁₋₄alkyl, piperazinyl, C₁₋₄alkylpiperazinyl, C₁₋₄alkylpiperazinylC₁₋₄alkyloxy, piperazinylC₁₋₄alkyl, C₁₋₄alkylpiperazinylC₁₋₄alkyl, C₁₋₄alkylpiperazinylC₁₋₄alkylamino, C₁₋₄alkylpiperazinylC₁₋₄alkylaminoC₁₋₆alkyl, tetrahydropyrimidinylpiperazinyl, tetrahydropyrimidinylpiperazinylC₁₋₄alkyl, piperidinylaminoC₁₋₄alkylamino, piperidinylaminoC₁₋₄alkylaminoC₁₋₄alkyl, (C₁₋₄alkylpiperidinyl)(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkylamino, (C1_alkylpiperidinyl)(hydroxyC1_alkyl)aminoC1_alkylaminoC1_alkyl, pyridinylC₁₋₄alkyloxy, hydroxyC₁₋₄alkylamino, hydroxyC₁₋₄alkylaminoC₁₋₄alkyl. di(C_{1.4}alkyl)aminoC_{1.4}alkylamino, aminothiadiazolyl, aminosulfonylpiperazinylC₁₋₄alkyloxy, or thiophenylC₁₋₄alkylamino; each R⁶ and R⁷ can be placed on the nitrogen in replacement of the hydrogen: aryl in the above is phenyl, or phenyl substituted with one or more substituents each independently selected from halo, C1-6alkyl, C1-6alkyloxy, trifluoromethyl, cyano or hydroxycarbonyl.

202. The compound of claim 201 wherein each of R², R³, R⁴ and R⁵ corresponds to R¹², R¹³, R¹⁴, and R¹⁵, respectively, in claim 201 wherein:

n is 0, 1 or 2;
t is 0, 1, 2 or 3;

$$O$$
 is O , O O , O O O

 R^2 is hydrogen, C_{1-6} alkyl or naphtalenylsulfonylpyrazinyl; each R^3 independently represents a hydrogen atom; R^4 is hydrogen, hydroxy, hydroxy C_{1-6} alkyl or C_{1-6} alkyloxy; R^5 is hydrogen, C_{1-6} alkyl, hydroxy C_{1-6} alkyl or C_{1-6} alkyloxy C_{1-6} alkyl;

is a radical selected from (a-1), (a-7) or (a-20);

each s is independently 0 or 1;

each R^6 is independently selected from hydrogen; thiophenyl; furanyl; benzofuranyl; phenyl; or phenyl substituted with one substituents independently selected from C_{1-6} alkyl, C_{1-6} alkyloxy, hydroxy C_{1-4} alkyl, C_{1-4} alkylsulfonyl or di $(C_{1-4}$ alkyl)amino; each R^7 is independently selected from hydrogen.

203. The compound of claim 201 wherein each of R², R³, R⁴ and R⁵ corresponds to R¹², R¹³, R¹⁴, and R¹⁵, respectively, claim 201 wherein:

n is 1 or 2;

$$\frac{1 \text{ t is } 0, 1, 2 \text{ or } 3;}{\text{Q is}}$$
, $-\text{CR} \left(\text{, or } -\text{CH} \right)$;

R² is hydrogen or C₁₋₆alkyl;

each R³ independently represents a hydrogen atom;

R⁴ is hydrogen;

R⁵ is hydrogen or C₁₋₆alkyloxyC₁₋₆alkyl;

—A is a radic

) is a radical selected from (a-1) or (a-20);

each s is independently 0 or 1;

each R^6 is independently selected from hydrogen; thiophenyl; furanyl; benzofuranyl; phenyl; or phenyl substituted with one substituents independently selected from C_{1-6} alkyl, C_{1-6} alkyloxy, hydroxy C_{1-4} alkyl or di(C_{1-4} alkyl)amino.

204. The compound of claim 201 wherein R¹² is H.

205. The compound of claim 201 wherein each of R², R³, R⁴ and R⁵ corresponds to R¹², R¹³, R¹⁴, and R¹⁵, respectively, in claim 201 wherein: t is 0;

- R² is hydrogen, halo, hydroxy, amino, nitro, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl or di(C₁₋₆alkyl)amino;
- R⁴ is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyloxy, arylC₁₋₆alkyl, aminoCarbonyl, aminoC₁₋₆alkyl, C₁₋₆alkylaminoC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl;

R⁵ is hydrogen

is a radical selected from (a-1), (a-3), (a-4), (a-5), (a-6), (a-7), (a-8), (a-9), (a-10), (a-11), (a-12), (a-13), (a-14), (a-15), (a-16), (a-17), (a-18), (a-19), (a-20), (a-21), (a-22), (a-23), (a-24), (a-25), (a-26), (a-28), (a-29), (a-30), (a-31), (a-32), (a-33), (a-34), (a-35), (a-36), (a-37), (a-38), (a-39), (a-40), (a-41), (a-42), (a-44), (a-45), (a-46), (a-47), (a-48) or (a-51); each s is independently 0, 1, 2, 3 or 4;

R⁶ is hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkylcxy; C₁₋₆alkylcarbonyl; C₁₋₆alkylcxycarbonyl; C₁₋₆alkylsulfonyl; hydroxyC₁₋₆alkyl; aryloxy; di(C₁₋₆alkyl)amino; cyano; thiophenyl; furanyl; furanyl substituted with hydroxyC₁₋₆alkyl; benzofuranyl; imidazolyl; oxazolyl; oxazolyl substituted with aryl and C₁₋₆alkyl; C₁₋₆alkyltriazolyl; tetrazolyl; pyrrolidinyl; pyrrolyl; morpholinyl; C₁₋₆alkylmorpholinyl; piperazinyl; C₁₋₆alkylpiperazinyl; hydroxyC₁₋₆alkylpiperazinyl; C₁₋₆alkyloxypiperidinyl; pyrazoly; pyrazolyl substituted with one or two substituents selected from C₁₋₆alkyl or trihaloC₁₋₆alkyl; pyridinyl; pyridinyl substituted with C₁₋₆alkyloxy, aryloxy or aryl; pyrimidinyl; quinolinyl; indole; phenyl; or phenyl substituted with one or two substituents independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy or trifluoromethyl;

 R^7 is hydrogen; halo; hydroxy; amino; nitro; trihalo C_{1-6} alkyl; trihalo C_{1-6} alkyloxy; C_{1-6} a

 C_{1-6} alkylsulfonyl; hydroxy C_{1-6} alkyl; aryloxy; di $(C_{1-6}$ alkyl)amino; cyano; pyridinyl; phenyl; or phenyl substituted with one or two substituents independently selected from halo, C_{1-6} alkyl, C_{1-6} alkyloxy or trifluoromethyl.

206. The compound of claim 201 wherein each of R², R³, R⁴ and R⁵ corresponds to R¹², R¹³, R¹⁴, and R¹⁵, respectively, in claim 201 wherein:

R⁵ is hydrogen, C₁₋₆alkyl, C₃₋₁₀cycloalkyl, hydroxyC₁₋₆alkyl, C₁₋₆alkyloxyC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl;

(a-9), (a-10), (a-11), (a-12), (a-13), (a-14), (a-15), (a-16), (a-17), (a-18), (a-20), (a-21), (a-22), (a-23), (a-24), (a-25), (a-26), (a-27), (a-28), (a-29), (a-31), (a-32), (a-34), (a-35), (a-36), (a-37), (a-38), (a-39), (a-40), (a-41), (a-42) (a-43) or (a-44);

l) each R⁶ and R⁷ are independently selected from hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkyl; C₁₋₆alkyl; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkylsulfonyl; cyanoC₁₋₆alkyl; hydroxyC₁₋₆alkyloxy; hydroxyC₁₋₆alkylamino; aminoC₁₋₆alkyloxy; di(C₁₋₆alkyl)aminocarbonyl; di(hydroxyC₁₋₆alkyl)amino; arylC₁₋₆alkyl)amino; di(C₁₋₆alkyl)aminoC₁₋₆alkyloxy; di(C₁₋₆alkyl)aminoC₁₋₆alkyloxy; di(C₁₋₆alkyl)aminoC₁₋₆alkyloxy; arylC₂₋₆alkenediyl; di(C₁₋₆alkyl)amino; di(C₁₋₆alkyl)aminoC₁₋₆alkyl; di(C₁₋₆alkyl)aminoC₁₋₆alkyl)aminoC₁₋₆alkyl; cyano; thiophenyl; thiophenyl substituted with di(C₁₋₆alkyl)aminoC₁₋₆alkyl(C₁₋₆alkyl)aminoC₁₋₆alkyl, di(C₁₋₆alkyl)aminoC₁₋₆alkyl; furanyl;

imidazolyl; C_{1-6} alkyltriazolyl; tetrazolyl; pyrrolidinyl; piperidinyl C_{1-6} alkyloxy; morpholinyl; C_{1-6} alkylmorpholinyl; morpholinyl C_{1-6} alkylpiperazinyl; C_{1-6} alkylpiperazinyl C_{1-6} alkylpiperazinyl C_{1-6} alkylpiperazinylsulfonyl; aminosulfonylpiperazinyl C_{1-6} alkylpiperazinyl C_{1-6} alkylpiperazinyl; aminosulfonylpiperazinyl C_{1-6} alkyl; di(C_{1-6} alkyl)aminosulfonylpiperazinyl; di(C_{1-6} alkyl)aminosulfonylpiperazinyl; di(C_{1-6} alkyl)aminosulfonylpiperazinyl; hydroxy C_{1-6} alkylpiperazinyl C_{1-6} alkyl; C_{1-6} alkyloxypiperidinyl; C_{1-6} alkyloxypiperidinyl C_{1-6} alkyl; hydroxy C_{1-6} alkylpiperazinyl; hydroxy C_{1-6} alkyloxy C_{1-6} alkylpiperazinyl C_{1-6} al

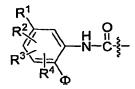
(hydroxyC₁₋₆alkyl)(C₁₋₆alkyl)amino; (hydroxyC₁₋₆alkyl)(C₁₋₆alkyl)aminoC₁₋₆alkyl; pyrrolidinylC₁₋₆alkyloxy; pyrazolyl; thiopyrazolyl; pyrazolyl substituted with two substituents selected from C₁₋₆alkyl or trihaloC₁₋₆alkyl; pyridinyl; pyridinyl substituted with C₁₋₆alkyloxy or aryl; pyrimidinyl; quinolinyl; indole; phenyl; phenyl substituted with one, two or three substituents independently selected from halo, amino, C₁₋₆alkyl, C₁₋₆alkyloxy, hydroxyC₁₋₄alkyl, trifluoromethyl, trifluoromethyloxy, hydroxyC₁₋₄alkyloxy, C₁₋₄alkyloxyC₁₋₄alkyloxy, aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)aminoC₁₋₄alkyl)aminoC₁₋₄alkyl, piperidinylC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, aminosulfonylpiperazinyl, aminosulfonylpiperazinylC₁₋₄alkyl, di(C₁₋₄alkyl)aminosulfonylpiperazinyl, di(C₁₋₄alkyl)aminosulfonylpiperazinyl, hydroxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, C₁₋₄alkyloxypiperidinyl, C₁₋₄alkylpiperazinylC₁₋₄alkyl, hydroxyC₁₋₄alkylpiperazinyl,

hydroxy C_{1-4} alkyloxy C_{1-4} alkylpiperazinyl C_{1-4} alkyl, (hydroxy C_{1-4} alkyl)(C_{1-4} alkyl)amino, (hydroxy C_{1-4} alkyl)(C_{1-4} alkyl)amino C_{1-4} alkyl, pyrrolidinyl C_{1-4} alkyloxy, morpholinyl C_{1-4} alkylpiperazinyl, C_{1-4} alkylpiperazinyl, C_{1-4} alkylpiperazinyl C_{1-4} alkylpiperazinyl, C_{1-4} alkylpiperazinyl C_{1-4} alkyloxy,

C₁₋₄alkylpiperazinylC₁₋₄alkyl, hydroxyC₁₋₄alkylamino, di(hydroxyC₁₋₄alkyl)amino, di(C₁₋₄alkyl)aminoC₁₋₄alkylamino, aminothiadiazolyl, aminosulfonylpiperazinylC₁₋₄alkyloxy, or thiophenylC₁₋₄alkylamino.

207. The compound of claim 201 that is selected from one of

wherein the terminal hydroxamic acid moiety (-C(O)-NH-OH) is replaced with



wherein Φ , R^1 , R^2 , R^3 , and R^4 are as defined in accordance with claim 1.

208. A compound according to claim 201 for use in inhibiting histone deacetylase.

209. A compound according to calim 201 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.

210. The compound of claim 209, wherein said treatment is effected by inhibiting histone deacetylase.

211. The compound of calim 209, wherein said cell proliferative disease is a neoplastic cell proliferative disease.

212. The compound of claim 209, wherein said cell proliferative disease is cancer.

213. The compound of claim 212, wherein said cancer is a solid tumor cancer.

214. The compound of claim 212, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.

215. A pharmaceutical composition comprising a compound according to claim 201 and a pharmaceutically acceptable carrier.

216. The pharmaceutical composition of claim 215 further comprising a nucleic acid level inhibitor of histone deacetylase.

217. The pharmaceutical composition of claim 216, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.

The pharmaceutical composition of claim 217, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.

219. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 201.

220. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 215.

- 221. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 216.
- 222. The method of claim 220, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 223. The method of claim 220, wherein said cell proliferative disease is cancer.
- 224. The method of claim 223, wherein said cancer is a solid tumor cancer.
- 225. The method of claim 224, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 226. The method of claim 221, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 227. The method of claim 221, wherein said cell proliferative disease is cancer.
- 228. The method of claim 227, wherein said cancer is a solid tumor cancer.
- 229. The method of claim 228, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 230. The compound of claim 201 wherein R2, R3, and R4 are all H.
- 231. A compound of the formula:

or a pharmaceutically acceptable salt thereof, wherein

 Φ is -NH $_2$ or -OH;

 R^1 is H or as defined in claim 1; R^2 . R^3 . and R^4 are as defined in claim 1;

n is 0, 1, 2 or 3 and when n is 0 then a direct bond is intended;

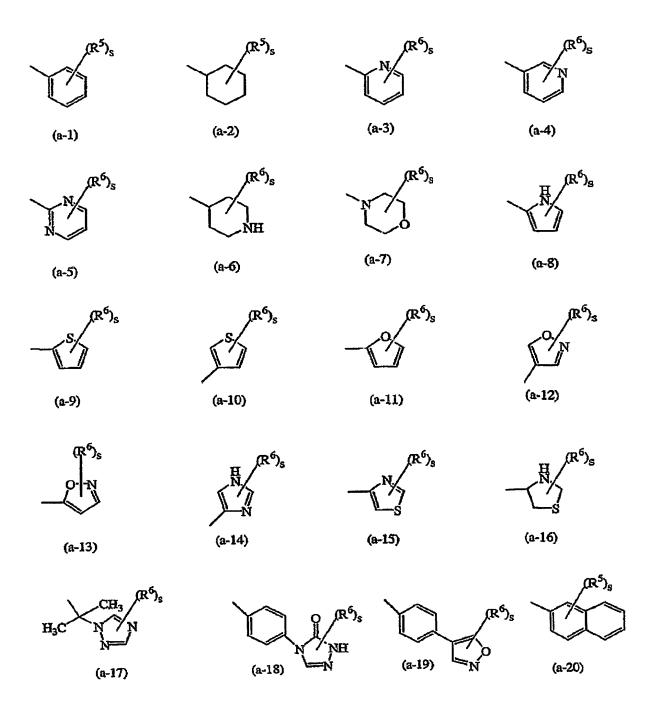
R is selected from the group consisting of hydrogen, halogen, -NH₂, nitro, hydroxy, aryl, heterocyclyl, C_3 - C_8 -cycloalkyl, heteroaryl, C_1 - C_7 -akyl, haloalkyl, C_1 - C_7 -alkenyl, C_1 - C_7 -alkyl-arylsulfanyl, C_1 - C_7 -alkyl-arylsulfinyl, C_1 - C_7 -alkyl-arylsulfonyl, C_1 - C_7 -alkyl-arylaminosulfonyl, C_1 - C_7 -alkyl-arylamine, C_1 - $C_$

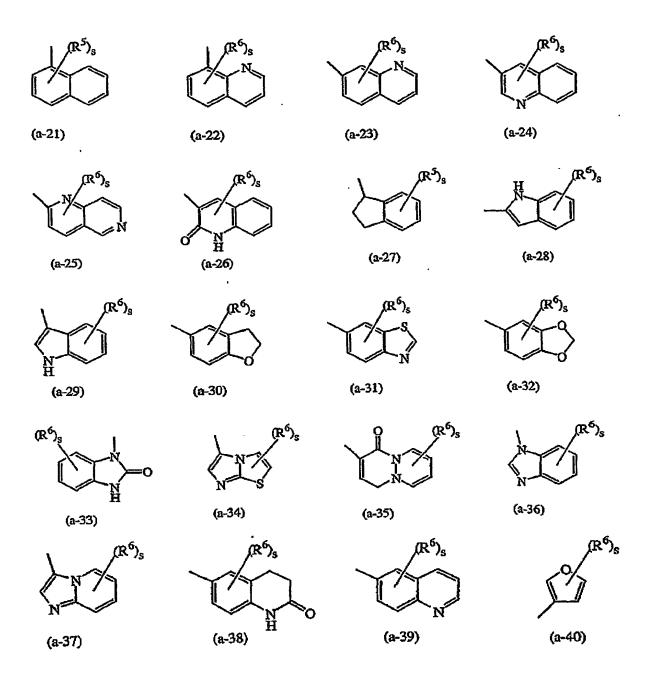
- R^{12} is hydrogen, halo, hydroxy, amino, nitro, C_{1-6} alkyl, C_{1-6} alkyloxy, trifluoromethyl, di(C_{1-6} alkyl)amino, hydroxyamino or naphtalenylsulfonylpyrazinyl;
- R¹³ is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyloxy, arylC₁₋₆alkyl, aminocarbonyl, hydroxycarbonyl, aminoC₁₋₆alkyl, aminocarbonylC₁₋₆alkyl, hydroxycarbonylC₁₋₆alkyl, hydroxyaminocarbonyl, C₁₋₆alkyloxycarbonyl, C₁₋₆alkylaminoC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl;

when Z is equal to nitrogen, then-L- is a direct bond;

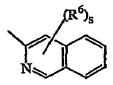
when Z is equal to __CH__, then _L_ is _NH_ or the bivalent radical _C1_6alkanediylNH_;

- R^{14} is hydrogen, C_{1-6} alkyl, C_{3-10} cycloalkyl, hydroxy C_{1-6} alkyl, C_{1-6} alkyloxy C_{1-6} alkyl, di $(C_{1-6}$ alkyl)amino C_{1-6} alkyl or aryl;
 - is a radical selected from

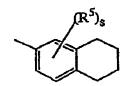




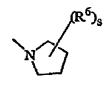
(a-41)



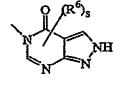
(a-42)



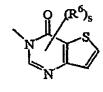
(a-43)



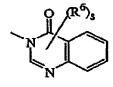
(a-44)



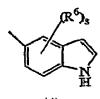
(a-45)



(a-46)

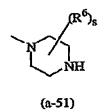


(a-47)



(a-48)

(a-50)



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wherein each s is independently 0, 1, 2, 3, 4 or 5;
each R<sup>5</sup> and R<sup>6</sup> are independently selected from hydrogen; halo; hydroxy; amino; nitro;
     trihaloC<sub>1-6</sub>alkyl; trihaloC<sub>1-6</sub>alkyloxy; C<sub>1-6</sub>alkyl; C<sub>1-6</sub>alkyl substituted with aryl and
     C<sub>3-10</sub>cycloalkyl; C<sub>1-6</sub>alkyloxy; C<sub>1-6</sub>alkyloxyC<sub>1-6</sub>alkyloxy; C<sub>1-6</sub>alkyloxy; C<sub>1-6</sub>alkylox
     C<sub>1-6</sub>alkyloxycarbonyl; C<sub>1-6</sub>alkylsulfonyl; cyanoC<sub>1-6</sub>alkyl; hydroxyC<sub>1-6</sub>alkyl;
     hydroxyC<sub>1-6</sub>alkyloxy; hydroxyC<sub>1-6</sub>alkylamino; aminoC<sub>1-6</sub>alkyloxy;
     di(C<sub>1-6</sub>alkyl)aminocarbonyl; di(hydroxyC<sub>1-6</sub>alkyl)amino; (aryl)(C<sub>1-6</sub>alkyl)amino;
     di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyloxy; di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkylamino;
     di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkylaminoC<sub>1-6</sub>alkyl; arylsulfonyl; arylsulfonylamino;
     aryloxy; aryloxyC<sub>1.6</sub>alkyl; arylC<sub>2.6</sub>alkenediyl; di(C<sub>1.6</sub>alkyl)amino;
     di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl; di(C<sub>1-6</sub>alkyl)amino(C<sub>1-6</sub>alkyl)amino;
     di(C_{1-6}alkyl)amino(C_{1-6}alkyl)amino(C_{1-6}alkyl)
     di(C_{1-6}alkyl)aminoC_{1-6}alkyl(C_{1-6}alkyl)amino;
     di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl;
     aminosulfonylamino(C1-6alkyl)amino;
     aminosulfonylamino(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl;
 di(C_{1-6}alkyl)aminosulfonylamino(C_{1-6}alkyl)amino;
 di(C_{1-6}alkyl)aminosulfonylamino(C_{1-6}alkyl)aminoC_{1-6}alkyl; cyano; thiophenyl;
thiophenyl substituted with di(C_{1-6}alkyl)aminoC_{1-6}alkyl(C_{1-6}alkyl)aminoC_{1-6}alkyl,
di(C_{1-6}alkyl)aminoC_{1-6}alkyl, C_{1-6}alkylpiperazinylC_{1-6}alkyl,
 hydroxyC<sub>1-6</sub>alkylpiperazinylC<sub>1-6</sub>alkyl,
 hydroxyC<sub>1-6</sub>alkyloxyC<sub>1-6</sub>alkylpiperazinylC<sub>1-6</sub>alkyl,
 di(C<sub>1-6</sub>alkyl)aminosulfonylpiperazinylC<sub>1-6</sub>alkyl.
C<sub>1-6</sub>alkyloxypiperidinyl, C<sub>1-6</sub>alkyloxypiperidinylC<sub>1-6</sub>alkyl, morpholinylC<sub>1-6</sub>alkyl,
hydroxyC_{1-6}alkyl(C_{1-6}alkyl)aminoC_{1-6}alkyl, or di(hydroxyC_{1-6}alkyl)aminoC_{1-6}alkyl;
furanyl; furanyl substituted with hydroxyC<sub>1.6</sub>alkyl; benzofuranyl; imidazolyl;
  oxazolyl; oxazolyl substituted with aryl and C_{1-6}alkyl; C_{1-6}alkyltriazolyl; tetrazolyl;
 pyrrolidinyl; pyrrolyl; piperidinylC<sub>1-6</sub>alkyloxy; morpholinyl; C<sub>1-6</sub>alkylmorpholinyl;
  morpholinylC<sub>1-6</sub>alkyloxy;
morpholinylC<sub>1-6</sub>alkyl; morpholinylC<sub>1-6</sub>alkylamino:
morpholinylC<sub>1-6</sub>alkylaminoC<sub>1-6</sub>alkyl; piperazinyl; C<sub>1-6</sub>alkylpiperazinyl;
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C₁₋₆ alkyIpiperazinylC ₁₋₈ alkyloxy**xy**; piperazinylC naphtalenylsulfonylpiperazinyl;naphtalenylsulfonylpiperidinyl;naphtalenylsulfony C_{1-6} alkyIpiperazinyl C_{1-6} alkyIpipera C₁_alkylpiperazinylC ₁_alkylaminoC ₁_alkylpiperazinylsulfonyl;[ony]; aminosulfonylpiperazinylC __alkyloxy;aminosulfonylpiperazinyl;azinyl; 1_alkyl;di(C 1_alkyl)aminosulfonylpiperazinyl; aminosulfonylpiperazinylC $\label{eq:dicondition} \mbox{di(C}_{1_a} \mbox{lkyl)} \ \ \mbox{aminosulfonylpiperazinylC} \qquad \mbox{$_{1_a}$lkyl;hydroxyC}_{1_a} \mbox{lkylpiperazinyl;} \ \ \mbox{inyl;}$ hydroxyC _alkylpiperazinylC _alkyl; C _alkyloxypiperidinyl yl; C₁_alkyloxypiperidinylC ₁_alkyl; piperidinylaminotoC₁_alkylamino; piperidinylamino10C _{1_a}lkylaminoC _{1_a}lkyl; (C₁_alkyl piperidinyl **yl**)(hydroxyC₁_alkyl)amino**1i**)C₁_alkylamino; **0**; (C_{1_alkylpiperidin} lyl)(hydroxyC _{1_alkyl})aminoC _{1_alkylpiperidin} lyl)(hydroxyC _{1_alkyl})aminoC _{1_alkylpiperidin} lyl)(hydroxyC _{1_alkylpiperidin} lyl)(hydrox hydroxyC ₁₋₈lkyloxyC ₁₋₆alkyIpiperazinyl; $\label{eq:hydroxyC} \ _{1\text{--}alkyloxyC} \ _{1\text{--}alkylpiperazinylC} \ _{1\text{--}alkyl} \text{ lkyl};$ (hydroxyC 1_alkyl)(C 1_alkyl)amino; ; (hydroxyC 1_alkyl)(C 1_alkyl)aminoC 1_alkyl; ; hydroxyC 1_alkylaminoC 1_alkyl; di(hydroxyC 1_alkyl)aminoC 1_alkyl; pyrrolidinylC ,_alkyl; pyrrolidinylC ,_alkyloxy; pyrazolyl; thiopyrazolyl; pyrazolyl substituted with two substituents selected from C_{1-a}lkyl or trihalo C_{1-a}lkyl; pyridinyl; pyridinyl substituted withC _{1...}alkyloxy, aryloxy or aryl; **p**pyrimidinyl; tetrahydropyrimdinylpiperazinyl; ; tetrahydropyrimdinylpiperazinylC , alkyl; kyl; quinolinyl; indolyl; phenyl; phenyl substituted with one, two or three substituents independently selected from halo, amino, nitro, C_{1-alkyl}, C_{1-alkyl}oxy, hydroxyC 1_alkyl, trifluoromethyl, trifluoromethyloxy, hydroxyC 1_alkyloxy,

C₁₋₄alkylsulfonyl, C₁₋₄alkyloxyC₁₋₄alkyloxy, C₁₋₄alkyloxy, di(C₁₋₄alkyl)aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)aminoC₁₋₄alkyl)aminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)aminoC₁₋₄alkyl)aminoC₁₋₄alkyl, aminosulfonylamino(C₁₋₄alkyl)amino, aminosulfonylamino(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminosulfonylamino(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)aminosulfonylamino(C₁₋₄alkyl)aminoC₁₋₆alkyl, cyano, piperidinylC₁₋₄alkyloxy, pyrrolidinylC₁₋₄alkyloxy, aminosulfonylpiperazinyl, aminosulfonylpiperazinylC₁₋₄alkyl, di(C₁₋₄alkyl)aminosulfonylpiperazinyl, di(C₁₋₄alkyl)aminosulfonylpiperazinylC₁₋₄alkyl, hydroxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, C₁₋₄alkyloxypiperidinyl, hydroxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, C₁₋₄alkyloxypiperidinyl,

C₁₋₄alkyloxypiperidinylC₁₋₄alkyl, hydroxyC₁₋₄alkyloxyC₁₋₄alkylpiperazinyl, hydroxyC₁₋₄alkyloxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, (hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)amino, (hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(hydroxyC₁₋₄alkyl)amino, di(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkyl, furanyl, furanyl substituted with -CH=CH-CH=CH-, pyrrolidinylC1-4alkyl, pyrrolidinylC1-4alkyloxy, morpholinyl, morpholinylC1_4alkyloxy, morpholinylC1_4alkyl, morpholinylC₁₋₄alkylamino, morpholinylC₁₋₄alkylaminoC₁₋₄alkyl, piperazinyl, C₁₋₄alkylpiperazinyl, C₁₋₄alkylpiperazinylC₁₋₄alkyloxy, piperazinylC₁₋₄alkyl, C₁₋₄alkylpiperazinylC₁₋₄alkyl, C₁₋₄alkylpiperazinylC₁₋₄alkylamino, C_{1-4} alkylpiperazinyl C_{1-4} alkylamino C_{1-6} alkyl, tetrahydropyrimidinylpiperazinyl, tetrahydropyrimidinylpiperazinylC₁₋₄alkyl, piperidinylaminoC₁₋₄alkylamino, piperidinylaminoC₁₋₄alkylaminoC₁₋₄alkyl, (C₁₋₄alkylpiperidinyl)(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkylamino, (C₁₋₄alkylpiperidinyl)(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkylaminoC₁₋₄alkyl, pyridinylC₁₋₄alkyloxy, hydroxyC₁₋₄alkylamino, hydroxyC₁₋₄alkylaminoC₁₋₄alkyl, di(C1-4alkyl)aminoC1-4alkylamino, aminothiadiazolyl, aminosulfonylpiperazinylC₁₋₄alkyloxy, or thiophenylC₁₋₄alkylamino; each R⁵ and R⁶ can be placed on the nitrogen in replacement of the hydrogen;

aryl in the above is phenyl, or phenyl substituted with one or more substituents each independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl, cyano or hydroxycarbonyl.

232. The compound of claim 231 wherein each of R^2 , R^3 , and R^4 corresponds to R^{12} , R^{13} , and R^{14} , respectively, in claim 231 wherein:

n is 1;
$$Q is - CR = CR = CH = CH$$

R² is hydrogen or nitro; R³ is hydrogen; when Z is equal to -CH, then -L- is the bivalent radical -C₁₋₆alkanediylNH-; R⁴ is hydrogen, C₁₋₆alkyl or aryl; is a radical selected from (a-1) or (a-21); each s is independently 0, 1 or 2; each R⁵ is independently selected from hydrogen; halo; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkyl; C₁₋₆alkyloxy; C₁₋₆alkylcarbonyl; aryloxy; cyano or phenyl. The compound of claim 231 wherein each of R², R³, and R⁴ corresponds to R¹², R¹³, and 233. R14, respectively, in claim 231 wherein: n is 1; -c, -cR, -cH, cH, ceach X is nitrogen; each Y is nitrogen; R² is hydrogen; R³ is hydrogen; when Z is equal to -CH, then -L is the bivalent radical -C₁₋₆alkanediylNH-; R4 is hydrogen, C1-6alkyl or aryl; is the radical (a-1); each s is independently 0 or 1; each R⁵ is independently selected from hydrogen or phenyl. The compound of claim 231 wherein each of R², R³, and R⁴ corresponds to R¹², R¹³, and 234. R¹⁴, respectively, in claim 231 wherein: each Z is N;

 R^2 is hydrogen, halo, hydroxy, amino, nitro, C_{1-6} alkyloxy, trifluoromethyl or $di(C_{1-6}$ alkyl) amino;

 R^3 is hydrogen, hydroxy, amino, hydroxy C_{1-6} alkyl, C_{1-6} alkyl, C_{1-6} alkyloxy, aryl C_{1-6} alkyl, aminocarbonyl, amino C_{1-6} alkyl, C_{1-6} alkylamino C_{1-6} alkyl; or di $(C_{1-6}$ alkyl)amino C_{1-6} alkyl;

R4 is hydrogen;

is a radical selected from (a-1), (a-3), (a-4), (a-5), (a-6), (a-7), (a-8), (a-9), (a-10), (a-11), (a-12), (a-13), (a-14), (a-15), (a-16), (a-17), (a-18), (a-19), (a-20), (a-21), (a-22), (a-23), (a-24), (a-25), (a-26), (a-28), (a-29), (a-30), (a-31), (a-32), (a-33), (a-34), (a-35), (a-36), (a-37), (a-38), (a-39), (a-40), (a-41), (a-42), (a-44), (a-45), (a-46), (a-47), (a-48) or (a-51);

each s is independently 0, 1, 2, 3 or 4;

R⁵ is hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkyl; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkylsulfonyl; hydroxyC₁₋₆alkyl; aryloxy; di(C₁₋₆alkyl)amino; cyano; thiophenyl; furanyl; substituted with hydroxyC₁₋₆alkyl; benzofuranyl; imidazolyl; oxazolyl; oxazolyl substituted with aryl and C₁₋₆alkyl; C₁₋₆alkyltriazolyl; tetrazolyl; pyrrolidinyl; pyrrolyl; morpholinyl; C₁₋₆alkylmorpholinyl; piperazinyl; C₁₋₆alkylpiperazinyl;

hydroxyC₁₋₆alkylpiperazinyl; C₁₋₆alkyloxypiperidinyl; pyrazolyl; pyrazolyl substituted with one or two substituents selected from C₁₋₆alkyl or trihaloC₁₋₆alkyl; pyridinyl; pyridinyl substituted with C₁₋₆alkyloxy, aryloxy or aryl; pyrimidinyl; quinolinyl; indole; phenyl; or phenyl substituted with one or two substituents independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy or trifluoromethyl; R⁶ is hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkylcarbonyl; C₁₋₆alkyloxycarbonyl; C₁₋₆alkylsulfonyl; hydroxyC₁₋₆alkyl; aryloxy; di(C₁₋₆alkyl)amino; cyano; pyridinyl; phenyl; or phenyl substituted with one or two substituents independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy or trifluoromethyl.

235. The compound of claim 231 that is selected from one of

wherein the terminal hydroxamic acid moiety (-C(O)-NH-OH) is replaced with

wherein Φ, R¹, R², R³, and R⁴ are as defined in accordance with claim 1.

236. The compound of claim 231 wherein R1, R2, R3, and R4 are all H.

237. A compound according to claim 231 for use in inhibting histone deacetylase.

238. A compound according to calim 231 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.

239. The compound of claim 238, wherein said treatment is effected by inhibiting histone deacetylase.

240. The compound of calim 238, wherein said cell proliferative disease is a neoplastic cell proliferative disease.

241. The compound of claim 238, wherein said cell proliferative disease is cancer.

242. The compound of claim 241, wherein said cancer is a solid tumor cancer.

243. The compound of claim 241, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.

244. A pharmaceutical composition comprising a compound according to claim 231 and a pharmaceutically acceptable carrier.

245. The pharmaceutical composition of claim 244 further comprising a nucleic acid level inhibitor of histone deacetylase.

246. The pharmaceutical composition of claim 245, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.

The pharmaceutical composition of claim 246, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.

A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 231.

A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 244.

- 250. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 245.
- 251. The method of claim 249, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 252. The method of claim 249, wherein said cell proliferative disease is cancer.
- 253. The method of claim 252, wherein said cancer is a solid tumor cancer.
- 254. The method of claim 253, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 255. The method of claim 250, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 256. The method of claim 250, wherein said cell proliferative disease is cancer.
- 257. The method of claim 256, wherein said cancer is a solid tumor cancer.
- 258. The method of claim 257, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 259. A compound of the formula:

or a pharmaceutically acceptable salt thereof, wherein

O is -NH 2 or -OH;

R1 is H or as defined in claim 1

R², R³, and R⁴ are as defined in claim 1;

n is 0, 1, 2 or 3 and when n is 0 then a direct bond is intended; m is 0, 1, 2 or 3 and when m is 0 then a direct bond is intended; t is 0 or 1 and when t is 0 then a direct bond is intended;

Q is nitrogen or —CK, or —CK;

X is nitrogen or —CK;

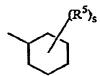
Y is nitrogen or —CK;

Z is -CH₂- or -O-;

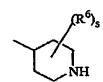
- R is selected from the group consisting of hydrogen, halogen, -NH₂, nitro, hydroxy, aryl, heterocyclyl, C_3 - C_8 -cycloalkyl, heteroaryl, C_1 - C_7 -akyl, haloalkyl, C_1 - C_7 -alkenyl, C_1 - C_7 -alkyl-arylsulfanyl, C_1 - C_7 -alkyl-arylsulfanyl, C_1 - C_7 -alkyl-arylsulfonyl, C_1 - C_7 -alkyl-arylsulfonyl, C_1 - C_7 -alkyl-arylamine, C_1 - C_1 -
- R¹² is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyloxy, arylC₁₋₆alkyl, aminocarbonyl, hydroxycarbonyl, aminoC₁₋₆alkyl, aminocarbonylC₁₋₆alkyl, hydroxycarbonylC₁₋₆alkyl, hydroxyaminocarbonyl, C₁₋₆alkyloxycarbonyl, C₁₋₆alkylaminoC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl;
 - -L- is a bivalent radical selected from C_{1-6} alkanediyl, carbonyl, sulfonyl, or C_{1-6} alkanediyl substituted with phenyl;
 - A is a radical selected from



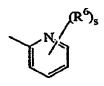
(a-5)



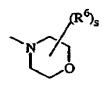
(a-2)



(a-6)

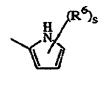


(a-3)



(a-7)

(a-4)



(a-8)

wherein each s is independently 0, 1, 2, 3, 4 or 5;

each R⁵ and R⁶ are independently selected from hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkyl; C₁₋₆alkyl substituted with aryl and C₃₋₁₀cycloalkyl; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkyloxyC₁₋₆alkyl; hydroxyC₁₋₆alkyl;

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hydroxyC<sub>1-6</sub>alkyloxy; hydroxyC<sub>1-6</sub>alkylamino; aminoC<sub>1-6</sub>alkyloxy;
di(C<sub>1.6</sub>alkyl)aminocarbonyl; di(hydroxyC<sub>1.6</sub>alkyl)amino; (aryl)(C<sub>1.6</sub>alkyl)amino;
di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyloxy; di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkylamino;
di(C_{1-6}alkyl)aminoC_{1-6}alkylaminoC_{1-6}alkyl; arylsulfonyl; arylsulfonylamino;
aryloxy; aryloxyC<sub>1-6</sub>alkyl; arylC<sub>2-6</sub>alkenediyl; di(C<sub>1-6</sub>alkyl)amino;
di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl; di(C<sub>1-6</sub>alkyl)amino(C<sub>1-6</sub>alkyl)amino;
di(C<sub>1-6</sub>alkyl)amino(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl;
di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl(C<sub>1-6</sub>alkyl)amino;
di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl;
aminosulfonylamino(C<sub>1.6</sub>alkyl)amino;
aminosulfonylamino(C1-6alkyl)aminoC1-6alkyl;
di(C<sub>1-6</sub>alkyl)aminosulfonylamino(C<sub>1-6</sub>alkyl)amino;
di(C<sub>1-6</sub>alkyl)aminosulfonylamino(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl; cyano; thiophenyl;
thiophenyl substituted with di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl,
di(C1_6alkyl)aminoC1_6alkyl, C1_6alkylpiperazinylC1_6alkyl,
hydroxyC<sub>1-6</sub>alkylpiperazinylC<sub>1-6</sub>alkyl,
hydroxyC<sub>1-6</sub>alkyloxyC<sub>1-6</sub>alkylpiperazinylC<sub>1-6</sub>alkyl,
di(C<sub>1-6</sub>alkyl)aminosulfonylpiperazinylC<sub>1-6</sub>alkyl,
C<sub>1-6</sub>alkyloxypiperidinyl, C<sub>1-6</sub>alkyloxypiperidinylC<sub>1-6</sub>alkyl, morpholinylC<sub>1-6</sub>alkyl,
hydroxyC_{1-6}alkyl(C_{1-6}alkyl)aminoC_{1-6}alkyl, or di(hydroxyC_{1-6}alkyl)aminoC_{1-6}alkyl;
furanyl; furanyl substituted with hydroxyC<sub>1-6</sub>alkyl; benzofuranyl; imidazolyl;
oxazolyl; oxazolyl substituted with aryl and C1-6alkyl; C1-6alkyltriazolyl; tetrazolyl;
pyrrolidinyl; pyrrolyl; piperidinylC<sub>1-6</sub>alkyloxy; morpholinyl; C<sub>1-6</sub>alkylmorpholinyl;
morpholinylC<sub>1-6</sub>alkyloxy;
morpholinylC<sub>1-6</sub>alkyl; morpholinylC<sub>1-6</sub>alkylamino;
morpholinylC<sub>1-6</sub>alkylaminoC<sub>1-6</sub>alkyl; piperazinyl; C<sub>1-6</sub>alkylpiperazinyl;
C<sub>1-6</sub>alkylpiperazinylC<sub>1-6</sub>alkyloxy; piperazinylC<sub>1-6</sub>alkyl;
naphtalenylsulfonylpiperazinyl; naphtalenylsulfonylpiperidinyl; naphtalenylsulfonyl;
C_{1-6}alkylpiperazinylC_{1-6}alkyl; C_{1-6}alkylpiperazinylC_{1-6}alkylamino;
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 $\label{eq:c1-6alkylpiperazinylC1-6alkylaminoC1-6alkyl; C1-6alkylpiperazinylsulfonyl; aminosulfonylpiperazinylC1-6alkyloxy; aminosulfonylpiperazinyl; aminosulfonylpiperazinylC1-6alkyl; di(C1-6alkyl)aminosulfonylpiperazinyl; di(C1-6alkyl)aminosulfonylpiperazinylC1-6alkyl; hydroxyC1-6alkylpiperazinyl; hydroxyC1-6alkylpiperazinylC1-6alkyl; C1-6alkyloxypiperidinyl; C1-6alkyloxypiperidinylC1-6alkyl; piperidinylaminoC1-6alkylamino; piperidinylaminoC1-6alkyl; (C1-6alkylpiperidinyl)(hydroxyC1-6alkyl)aminoC1-6alkylamino; (C1-6alkylpiperidinyl)(hydroxyC1-6alkyl)aminoC1-6alkylaminoC1-6alkyl; hydroxyC1-6alkyloxyC1-6alkylpiperazinyl; hydroxyC1-6alkyloxyC1-6alkylpiperazinylC1-6alkyl; (hydroxyC1-6alkyl)(C1-6alkyl)aminoC1-6alkyl; hydroxyC1-6alkyl)(C1-6alkyl)aminoC1-6alkyl; hydroxyC1-6alkylaminoC1-6alkyl; di(hydroxyC1-6alkyl)aminoC1-6alkyl;$

pyrrolidinylC₁₋₆alkyl; pyrrolidinylC₁₋₆alkyloxy; pyrazolyl; thiopyrazolyl; pyrazolyl substituted with two substituents selected from C₁₋₆alkyl or trihaloC₁₋₆alkyl; pyridinyl; pyridinyl substituted with C₁₋₆alkyloxy, aryloxy or aryl; pyrimidinyl; tetrahydropyrimidinylpiperazinyl; tetrahydropyrimidinylpiperazinylC₁₋₆alkyl; quinolinyl; indolyl; phenyl; phenyl substituted with one, two or three substituents independently selected from halo, amino, nitro, C₁₋₆alkyl, C₁₋₆alkyloxy, hydroxyC₁₋₄alkyl, trifluoromethyl, trifluoromethyloxy, hydroxyC₁₋₄alkyloxy, C₁₋₄alkylsulfonyl, C₁₋₄alkyloxyC₁₋₄alkyloxy, C₁₋₄alkyloxycarbonyl, aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)aminocarbonyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)amino(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)amino(C₁₋₄alkyl)aminoC₁₋₄alkyl, $di(C_{1-4}alkyl)aminoC_{1-4}alkyl(C_{1-4}alkyl)amino,$ di(C₁₋₄alkyl)aminoC₁₋₄alkyl(C₁₋₄alkyl)aminoC₁₋₄alkyl, aminosulfonylamino(C1-4alkyl)amino, aminosulfonylamino(C1-4alkyl)aminoC1-4alkyl, di(C₁₋₄alkyl)aminosulfonylamino(C₁₋₄alkyl)amino,

di(C₁₋₄alkyl)aminosulfonylamino(C₁₋₄alkyl)aminoC₁₋₆alkyl, cyano, piperidinylC₁₋₄alkyloxy, pyrrolidinylC₁₋₄alkyloxy, aminosulfonylpiperazinyl, aminosulfonylpiperazinylC1-4alkyl, di(C1-4alkyl)aminosulfonylpiperazinyl, di(C₁₋₄alkyl)aminosulfonylpiperazinylC₁₋₄alkyl, hydroxyC₁₋₄alkylpiperazinyl, hydroxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, C₁₋₄alkyloxypiperidinyl, C₁₋₄alkyloxypiperidinylC₁₋₄alkyl, hydroxyC₁₋₄alkyloxyC₁₋₄alkylpiperazinyl, hydroxyC₁₋₄alkyloxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, (hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)amino, (hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(hydroxyC₁₋₄alkyl)amino, di(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkyl, furanyl, furanyl substituted with -CH=CH-CH=CH-, pyrrolidinylC₁₋₄alkyl, pyrrolidinylC₁₋₄alkyloxy, morpholinyl, morpholinylC₁₋₄alkyloxy, morpholinylC₁₋₄alkyl, morpholinylC₁₋₄alkylamino, morpholinylC₁₋₄alkylaminoC₁₋₄alkyl, piperazinyl, C₁₋₄alkylpiperazinyl, C₁₋₄alkylpiperazinylC₁₋₄alkyloxy, piperazinylC₁₋₄alkyl, C_{1.4}alkylpiperazinylC_{1.4}alkyl, C_{1.4}alkylpiperazinylC_{1.4}alkylamino, C_{1-4} alkylpiperazinyl C_{1-4} alkylamino C_{1-6} alkyl, tetrahydropyrimidinylpiperazinyl, tetrahydropyrimidinylpiperazinylC_{1.4}alkyl, piperidinylaminoC_{1.4}alkylamino, piperidinylaminoC₁₋₄alkylaminoC₁₋₄alkyl, (C1-4alkylpiperidinyl)(hydroxyC1-4alkyl)aminoC1-4alkylamino, (C₁₋₄alkylpiperidinyl)(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkylaminoC₁₋₄alkyl, pyridinylC₁₋₄alkyloxy,

hydroxyC₁₋₄alkylamino, hydroxyC₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkylamino, aminothiadiazolyl, aminosulfonylpiperazinylC₁₋₄alkyloxy, or thiophenylC₁₋₄alkylamino; each R⁵ and R⁶ can be placed on the nitrogen in replacement of the hydrogen;

aryl in the above is phenyl, or phenyl substituted with one or more substituents each independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy, trifluoromethyl, cyano or hydroxycarbonyl.

260. The compound of claim 259 wherein each of R², R³, and R⁴ corresponds to R¹², R¹³, and R¹⁴, respectively, in claim 259 wherein:

```
n is 0, 1 or 2;
m is 0, 1 or 2;
each Q is -C \leqslant;
each X is nitrogen;
R<sup>2</sup> is hydrogen:
-L- is a bivalent radical selected from carbonyl, sulfonyl, or C<sub>1-6</sub>alkanediyl
  substituted with phenyl;
             is a radical selected from (a-1), (a-20) or (a-43);
each s is independently 0 or 1;
each R<sup>5</sup> is independently selected from hydrogen or phenyl.
            The compound of claim 259 wherein each of R<sup>2</sup>, R<sup>3</sup>, and R<sup>4</sup> corresponds to R<sup>12</sup>, R<sup>13</sup>, and
261.
R<sup>14</sup>, respectively, in claim 259 wherein:
n is 0, 1 or 2;
m is 1 or 2;
       O is
       X is nitrogen;
R<sup>2</sup> is hydrogen;
-L- is a bivalent radical selected from carbonyl or sulfonyl;
             is a radical selected from (a-1) or (a-20);
each s is independently 0 or 1;
each R<sup>5</sup> is independently selected from hydrogen or aryl.
            The compound of claim 259 wherein each of R<sup>2</sup>, R<sup>3</sup>, and R<sup>4</sup> corresponds to R<sup>12</sup>, R<sup>13</sup>, and
262.
R<sup>14</sup>, respectively, in claim 259 wherein:
t is 0;
```

R² is hydrogen, hydroxy, amino, hydroxyC_{1.6}alkyl, C_{1.6}alkyl, C_{1.6}alkyloxy, arylC_{1.6}alkyl, aminocarbonyl, aminoC_{1.6}alkyl, C_{1.6}alkylaminoC_{1.6}alkyl or di(C_{1.6}alkyl)aminoC_{1.6}alkyl;

-L- is a bivalent radical selected from C₁₋₆alkanediyl, carbonyl or sulfonyl;

is a radical selected from (a-1), (a-3), (a-4), (a-5), (a-6), (a-7), (a-8), (a-9), (a-10), (a-11), (a-12), (a-13), (a-14), (a-15), (a-16), (a-17), (a-18), (a-19), (a-20), (a-21), (a-22), (a-23), (a-24), (a-25), (a-26), (a-28), (a-29), (a-30), (a-31), (a-32), (a-33), (a-34), (a-35), (a-36), (a-37), (a-38), (a-39), (a-40), (a-41), (a-42), (a-44), (a-45), (a-46), (a-47), (a-48) or (a-51);

each s is independently 0, 1, 2, 3 or 4;

R⁵ is hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkyl; C₁₋₆alkyloxy; C₁₋₆alkylcarbonyl; C₁₋₆alkyloxycarbonyl; C₁₋₆alkylsulfonyl; hydroxyC₁₋₆alkyl; aryloxy; di(C₁₋₆alkyl)amino; cyano; thiophenyl; furanyl substituted with hydroxyC₁₋₆alkyl; benzofuranyl; imidazolyl; oxazolyl; oxazolyl substituted with aryl and C₁₋₆alkyl; C₁₋₆alkyltriazolyl; tetrazolyl; pyrrolidinyl; pyrrolyl; morpholinyl;

C₁₋₆alkylmorpholinyl; piperazinyl; C₁₋₆alkylpiperazinyl; hydroxyC₁₋₆alkylpiperazinyl; C₁₋₆alkyloxypiperidinyl; pyrazoly; pyrazolyl substituted with one or two substituents selected from C₁₋₆alkyl or trihaloC₁₋₆alkyl; pyridinyl; pyridinyl substituted with C₁₋₆alkyloxy, aryloxy or aryl; pyrimidinyl; quinolinyl; indole; phenyl; or phenyl substituted with one or two substituents independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy or trifluoromethyl; R⁶ is hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkyloxy; C₁₋₆alkylcarbonyl; C₁₋₆alkyloxycarbonyl; C₁₋₆alkylsulfonyl; hydroxyC₁₋₆alkyl; aryloxy; di(C₁₋₆alkyl)amino; cyano; pyridinyl; phenyl; or phenyl substituted with one or two substituents independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy or trifluoromethyl.

263. The compound of claim 259 that is selected from one of

wherein the terminal hydroxamic acid moiety (-C(O)-NH-OH) is replaced with

wherein Φ , R^1 , R^2 , R^3 , and R^4 are as defined in accordance with claim 1.

264. The compound of claim 259 R1, R2, R3, and R4 are all H.

265. A compound according to claim 259 for use in inhibiting histone deacetylase.

266. A compound according to calim 259 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.

267. The compound of claim 266, wherein said treatment is effected by inhibiting histone deacetylase.

268. The compound of calim 266, wherein said cell proliferative disease is a neoplastic cell proliferative disease.

269. The compound of claim 266, wherein said cell proliferative disease is cancer.

270. The compound of claim 269, wherein said cancer is a solid tumor cancer.

271. The compound of claim 269, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.

272. A pharmaceutical composition comprising a compound according to claim 259 and a pharmaceutically acceptable carrier.

273. The pharmaceutical composition of claim 272 further comprising a nucleic acid level inhibitor of histone deacetylase.

274. The pharmaceutical composition of claim 273, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.

275. The pharmaceutical composition of claim 274, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.

276. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 259.

277. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 272.

- 278. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 273.
- 279. The method of claim 277, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 280. The method of claim 277, wherein said cell proliferative disease is cancer.
- 281. The method of claim 280, wherein said cancer is a solid tumor cancer.
- 282. The method of claim 281, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 283. The method of claim 278, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 284. The method of claim 278, wherein said cell proliferative disease is cancer.
- 285. The method of claim 284, wherein said cancer is a solid tumor cancer.
- 286. The method of claim 285, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 287. A compound of the formula:

$$R^{1}$$
 R^{2}
 R^{3}
 $= |=$
 R^{4}
 $Q = X$
 Z
 $(CH_{2})_{t} - L$
 A

or a pharmaceutically acceptable salt thereof, wherein

 Φ is -NH₂ or -OH;

R1 is H or as defined in claiml;

R², R³, and R⁴ are as defined in claim 1;

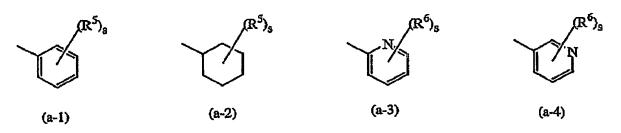
t is 0, 1, 2, 3 or 4 and when t is 0 then a direct bond is intended;

X is nitrogen or — ;

Y is nitrogen or —

Z is -NH-, -O- or -CH₂-;

- R is selected from the group consisting of hydrogen, halogen, -NH₂, nitro, hydroxy, aryl, heterocyclyl, C_3 - C_8 -cycloalkyl, heteroaryl, C_1 - C_7 -akyl, haloalkyl, C_1 - C_7 -alkenyl, C_1 - C_7 -alkyl-aryloxy, C_1 - C_7 -alkyl-arylsulfanyl, C_1 - C_7 -alkyl-arylsulfinyl, C_1 - C_7 -alkyl-arylaminosulfonyl, C_1 - C_7 -alkyl-arylamine, C_1 - C_7 -alkynyl-C(O)-amine, C_1 - C_7 -alkynyl-C(O)-ami
- R¹² is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyloxy, arylC₁₋₆alkyl, aminocarbonyl, hydroxycarbonyl, aminoC₁₋₆alkyl, aminocarbonylC₁₋₆alkyl, hydroxycarbonylC₁₋₆alkyl, hydroxyaminocarbonyl, C₁₋₆alkyloxycarbonyl, C₁₋₆alkylaminoC₁₋₆alkyl or di(C₁₋₆alkyl)aminoC₁₋₆alkyl;
 - -L- is a bivalent radical selected from -NR⁹C(O)-, -NR⁹SO₂- or -NR⁹CH₂-wherein R⁹ is hydrogen, C₁₋₆alkyl, C₃₋₁₀cycloalkyl, hydroxyC₁₋₆alkyl, C₁₋₆alkyl) aminoC₁₋₆alkyl;
 - is a radical selected from



$$(a-5) \qquad (a-6) \qquad (a-7) \qquad (a-8)$$

$$(a-7) \qquad (a-8)$$

$$(a-9) \qquad (a-10) \qquad (a-11) \qquad (a-12)$$

$$(a-12) \qquad (a-12)$$

$$(a-13) \qquad (a-14) \qquad (a-15) \qquad (a-16)$$

$$(a-17) \qquad (a-18) \qquad (a-19) \qquad (a-20)$$

$$(a-21) \qquad (a-22) \qquad (a-22) \qquad (a-24)$$

$$(a-25) \qquad (a-26) \qquad (a-27) \qquad (a-28)$$

wherein each s is independently 0, 1, 2, 3, 4 or 5; each R⁵ and R⁶ are independently selected from hydrogen; halo; hydroxy; amino; nitro;

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trihaloC<sub>1-6</sub>alkyl; trihaloC<sub>1-6</sub>alkyloxy; C<sub>1-6</sub>alkyl; C<sub>1-6</sub>alkyl substituted with aryl and
C_{3-10}cycloalkyl; C_{1-6}alkyloxy; C_{1-6}
C<sub>1-6</sub>alkyloxycarbonyl; C<sub>1-6</sub>alkylsulfonyl; cyanoC<sub>1-6</sub>alkyl; hydroxyC<sub>1-6</sub>alkyl;
hydroxyC<sub>1-6</sub>alkyloxy; hydroxyC<sub>1-6</sub>alkylamino; aminoC<sub>1-6</sub>alkyloxy;
di(C<sub>1-6</sub>alkyl)aminocarbonyl; di(hydroxyC<sub>1-6</sub>alkyl)amino; (aryl)(C<sub>1-6</sub>alkyl)amino;
di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyloxy; di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkylamino;
di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkylaminoC<sub>1-6</sub>alkyl; arylsulfonyl; arylsulfonylamino;
aryloxy; aryloxyC<sub>1-6</sub>alkyl; arylC<sub>2-6</sub>alkenediyl; di(C<sub>1-6</sub>alkyl)amino;
di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl; di(C<sub>1-6</sub>alkyl)amino(C<sub>1-6</sub>alkyl)amino;
di(C<sub>1-6</sub>alkyl)amino(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl;
di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl(C<sub>1-6</sub>alkyl)amino;
di(C_{1-6}alkyl)aminoC_{1-6}alkyl(C_{1-6}alkyl)aminoC_{1-6}alkyl;
aminosulfonylamino(C1-6alkyl)amino;
aminosulfonylamino(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl;
di(C<sub>1-6</sub>alkyl)aminosulfonylamino(C<sub>1-6</sub>alkyl)amino;
di(C<sub>1-6</sub>alkyl)aminosulfonylamino(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl; cyano; thiophenyl;
thiophenyl substituted with di(C1-6alkyl)aminoC1-6alkyl(C1-6alkyl)aminoC1-6alkyl,
di(C<sub>1-6</sub>alkyl)aminoC<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkylpiperazinylC<sub>1-6</sub>alkyl,
hydroxyC<sub>1-6</sub>alkylpiperazinylC<sub>1-6</sub>alkyl,
hydroxyC<sub>1-6</sub>alkyloxyC<sub>1-6</sub>alkylpiperazinylC<sub>1-6</sub>alkyl,
di(C<sub>1-6</sub>alkyl)aminosulfonylpiperazinylC<sub>1-6</sub>alkyl,
C<sub>1.6</sub>alkyloxypiperidinyl, C<sub>1.6</sub>alkyloxypiperidinylC<sub>1.6</sub>alkyl, morpholinylC<sub>1.6</sub>alkyl,
hydroxyC_{1-6}alkyl(C_{1-6}alkyl)aminoC_{1-6}alkyl, or di(hydroxyC_{1-6}alkyl)aminoC_{1-6}alkyl;
furanyl; furanyl substituted with hydroxyC<sub>1.6</sub>alkyl; benzofuranyl; imidazolyl;
oxazolyl; oxazolyl substituted with aryl and C_{1-6}alkyl; C_{1-6}alkyltriazolyl; tetrazolyl;
pyrrolidinyl; pyrrolyl; piperidinylC<sub>1-6</sub>alkyloxy; morpholinyl; C<sub>1-6</sub>alkylmorpholinyl;
morpholinylC<sub>1-6</sub>alkyloxy;
morpholinylC<sub>1-6</sub>alkyl; morpholinylC<sub>1-6</sub>alkylamino;
morpholinylC<sub>1.6</sub>alkylaminoC<sub>1.6</sub>alkyl; piperazinyl; C<sub>1.6</sub>alkylpiperazinyl;
 C<sub>1-6</sub>alkylpiperazinylC<sub>1-6</sub>alkyloxy; piperazinylC<sub>1-6</sub>alkyl;
 naphtalenylsulfonylpiperazinyl; naphtalenylsulfonylpiperidinyl; naphtalenylsulfonyl:
 C<sub>1-6</sub>alkylpiperazinylC<sub>1-6</sub>alkyl; C<sub>1-6</sub>alkylpiperazinylC<sub>1-6</sub>alkylamino;
 C_{1-6}alkylpiperazinylC_{1-6}alkylaminoC_{1-6}alkyl; C_{1-6}alkylpiperazinylsulfonyl;
 aminosulfonylpiperazinylC<sub>1-6</sub>alkyloxy; aminosulfonylpiperazinyl;
 aminosulfonylpiperazinylC<sub>1-6</sub>alkyl; di(C<sub>1-6</sub>alkyl)aminosulfonylpiperazinyl;
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di(C₁₋₆alkyl)aminosulfonylpiperazinylC₁₋₆alkyl; hydroxyC₁₋₆alkylpiperazinyl; hydroxyC₁₋₆alkylpiperazinylC₁₋₆alkyl; C₁₋₆alkyloxypiperidinyl; C₁₋₆alkyloxypiperidinylC₁₋₆alkyl; piperidinylaminoC₁₋₆alkylamino; piperidinylaminoC₁₋₆alkylaminoC₁₋₆alkyl; (C₁₋₆alkylpiperidinyl)(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkylamino; (C₁₋₆alkylpiperidinyl)(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkylaminoC₁₋₆alkyl; hydroxyC₁₋₆alkyloxyC₁₋₆alkylpiperazinyl; hydroxyC₁₋₆alkyloxyC₁₋₆alkylpiperazinylC₁₋₆alkyl; (hydroxyC₁₋₆alkyl)(C₁₋₆alkyl)amino; (hydroxyC₁₋₆alkyl)(C₁₋₆alkyl)aminoC₁₋₆alkyl; hydroxyC₁₋₆alkylaminoC₁₋₆alkyl; di(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkyl; pyrrolidinylC_{1.6}alkyl; pyrrolidinylC_{1.6}alkyloxy; pyrazolyl; thiopyrazolyl; pyrazolyl substituted with two substituents selected from C_{1.6}alkyl or trihaloC_{1.6}alkyl; pyridinyl; pyridinyl substituted with C₁₋₆alkyloxy, aryloxy or aryl; pyrimidinyl; tetrahydropyrimidinylpiperazinyl; tetrahydropyrimidinylpiperazinylC₁₋₆alkyl; quinolinyl; indole; phenyl; phenyl substituted with one, two or three substituents independently selected from halo, amino, nitro, C1-6alkyl, C1-6alkyloxy, hydroxyC₁₋₄alkyl, trifluoromethyl, trifluoromethyloxy, hydroxyC₁₋₄alkyloxy, C₁₋₄alkylsulfonyl, C₁₋₄alkyloxyC₁₋₄alkyloxy, C₁₋₄alkyloxycarbonyl, aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)aminocarbonyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, $di(C_{1-4}alkyl)aminoC_{1-4}alkylaminoC_{1-4}alkyl,$ di(C1-4alkyl)amino(C1-4alkyl)amino, di(C1-4alkyl)amino(C1-4alk di(C₁₋₄alkyl)aminoC₁₋₄alkyl(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)aminoC₁₋₄alkyl(C₁₋₄alkyl)aminoC₁₋₄alkyl, aminosulfonylamino(C1-4alkyl)amino, aminosulfonylamino(C1-4alkyl)aminoC1-4alkyl, di(C₁₋₄alkyl)aminosulfonylamino(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)aminosulfonylamino(C₁₋₄alkyl)aminoC₁₋₆alkyl, cyano, piperidinylC₁₋₄alkyloxy, pyrrolidinylC₁₋₄alkyloxy, aminosulfonylpiperazinyl, aminosulfonylpiperazinylC1-4alkyl, di(C1-4alkyl)aminosulfonylpiperazinyl, di(C₁₋₄alkyl)aminosulfonylpiperazinylC₁₋₄alkyl, hydroxyC₁₋₄alkylpiperazinyl, hydroxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, C₁₋₄alkyloxypiperidinyl, C₁₋₄alkyloxypiperidinylC₁₋₄alkyl, hydroxyC₁₋₄alkyloxyC₁₋₄alkylpiperazinyl, hydroxyC₁₋₄alkyloxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, (hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)amino, (hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)aminoC₁₋₄alkyl, di(hydroxyC₁₋₄alkyl)amino, di(hydroxyC₁₋₄alkyl)aminoC₁₋₄alkyl, furanyl, furanyl substituted with -CH=CH-CH=CH-, pyrrolidinylC₁₋₄alkyl, pyrrolidinylC₁₋₄alkyloxy, morpholinyl, morpholinylC₁₋₄alkyloxy, morpholinylC₁₋₄alkyl,

 $morpholinylC_{1-4}$ alkylamino, $morpholinylC_{1-4}$ alkylamino C_{1-4} alkyl, piperazinyl, C_{1-4} alkylpiperazinyl, C_{1-4} alkylpiperazinyl C_{1-4} alkyloxy, piperazinyl C_{1-4} alkyl, C_{1-4} alkylpiperazinyl C_{1-4} alkyl, C_{1-4} alkylpiperazinyl C_{1-4} alkylamino, C_{1-4} alkylpiperazinyl C_{1-4} alkylamino C_{1-6} alkyl, tetrahydropyrimidinylpiperazinyl, tetrahydropyrimidinylpiperazinylC₁₋₄alkyl, piperidinylaminoC₁₋₄alkylamino, piperidinylaminoC₁₋₄alkylaminoC₁₋₄alkyl, $(C_{1-4}alkylpiperidinyl)(hydroxyC_{1-4}alkyl)aminoC_{1-4}alkylamino,$ $(C_{1-4}alkylpiperidinyl)(hydroxyC_{1-4}alkyl)aminoC_{1-4}alkylaminoC_{1-4}alkyl,$ pyridinylC₁₋₄alkyloxy, hydroxyC₁₋₄alkylamino, hydroxyC₁₋₄alkylaminoC₁₋₄alkyl, $di(C_{1-4}alkyl)aminoC_{1-4}alkylamino, aminothiadiazolyl,$ aminosulfonylpiperazinyl C_{1-4} alkyloxy, or thiophenyl C_{1-4} alkylamino; each R⁵ and R⁶ can be placed on the nitrogen in replacement of the hydrogen; aryl in the above is phenyl, or phenyl substituted with one or more substituents each independently selected from halo, C1-6alkyl, C1-6alkyloxy, trifluoromethyl, cyano or hydroxycarbonyl.

288. The compound of claim 287 wherein each of R², R³, and R⁴ corresponds to R¹², R¹³, and R¹⁴, respectively, in claim 287 wherein:

t is 0 or 1;

$$Q$$
 is $-CR$, or $-CH$;

X is nitrogen;

R12 is hydrogen, hydroxy, C1.6alkyl, or arylC1.6alkyl;

-L- is a bivalent radical selected from -NHC(O)- or -NHSO₂-;

is a radical selected from (a-1) or (a-20);

each s is independently 0 or 1;

each R⁵ is independently selected from hydrogen or phenyl.

289. The compound of claim 287 wherein each of R², R³, and R⁴ corresponds to R¹², R¹³, and R¹⁴, respectively, in claim287 wherein:

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tis 1;

$$Q$$
 is $-CR$, or $-CH$;

X is nitrogen;

Y is nitrogen;

Z is -O- or -CH2-;

R12 is H;

-L- is a bivalent radical selected from -NHC(O)- or -NHSO₂-;



is a radical selected from (a-1) or (a-20);

each s is independently 0 or 1;

each R⁵ is independently selected from hydrogen or phenyl.

The compound of claim 287 wherein each of R², R³, and R⁴ corresponds to R¹², R¹³, and 290. R¹⁴, respectively, in claim 287 wherein:

t is 0:

R¹² is hydrogen, hydroxy, amino, hydroxyC₁₋₆alkyl, C₁₋₆alkyl, C₁₋₆alkyloxy,

 $arylC_{1-6}alkyl$, aminocarbonyl, amino $C_{1-6}alkyl$, $C_{1-6}alkyl$ amino $C_{1-6}alkyl$ or $di(C_{1-6}alkyl)aminoC_{1-6}alkyl;$

-L- is a bivalent radical selected from -NHC(O)- or -NHSO₂-;

is a radical selected from (a-1), (a-3), (a-4), (a-5), (a-6), (a-7), (a-8), (a-9), (a-10), (a-11), (a-12), (a-13), (a-14), (a-15), (a-16), (a-17), (a-18), (a-19), (a-20), (a-21), (a-22), (a-23), (a-24), (a-25), (a-26), (a-28), (a-29), (a-30), (a-31), (a-32), (a-33), (a-34), (a-35), (a-36), (a-37), (a-38), (a-39), (a-40), (a-41), (a-42), (a-44), (a-45), (a-46), (a-47), (a-48) or (a-51);

each s is independently 0, 1, 2, 3 or 4;

R⁵ is hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkyl; C₁₋₆alkyloxy; C₁₋₆alkylcarbonyl; C₁₋₆alkyloxycarbonyl; C₁₋₆alkylsulfonyl; hydroxyC₁₋₆alkyl; aryloxy; di(C₁₋₆alkyl)amino; cyano; thiophenyl; furanyl; substituted with hydroxyC₁₋₆alkyl; benzofuranyl; imidazolyl; oxazolyl; oxazolyl substituted with aryl and C₁₋₆alkyl; C₁₋₆alkyltriazolyl; tetrazolyl; pyrrolidinyl; pyrrolyl; morpholinyl; C₁₋₆alkylmorpholinyl; piperazinyl;

C₁₋₆alkylpiperazinyl; hydroxyC₁₋₆alkylpiperazinyl;

 $C_{1\text{-}6}$ alkyloxypiperidinyl; pyrazoly; pyrazolyl substituted with one or two substituents selected from $C_{1\text{-}6}$ alkyl or trihalo $C_{1\text{-}6}$ alkyl; pyridinyl; pyridinyl substituted with $C_{1\text{-}6}$ alkyloxy, aryloxy or aryl; pyrimidinyl; quinolinyl; indole; phenyl; or phenyl substituted with one or two substituents independently selected from halo, $C_{1\text{-}6}$ alkyl, $C_{1\text{-}6}$ alkyloxy or trifluoromethyl;

R⁶ is hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkyl; C₁₋₆alkyloxy; C₁₋₆alkylcarbonyl; C₁₋₆alkyloxycarbonyl; C₁₋₆alkylsulfonyl; hydroxyC₁₋₆alkyl; aryloxy; di(C₁₋₆alkyl)amino; cyano; pyridinyl; phenyl; or phenyl substituted with one or two substituents independently selected from halo, C₁₋₆alkyl, C₁₋₆alkyloxy or trifluoromethyl.

291. The compound of claim 287 wherein each of R², R³, and R⁴ corresponds to R¹², R¹³, and R¹⁴, respectively, in claim 287 wherein:

R³ and R⁴ are each independently selected from hydrogen, hydroxy, hydroxyC₁₋₆alkyl, aminoC₁₋₆alkyl or aminoaryl;

(a-9), (a-10), (a-11), (a-12), (a-13), (a-14), (a-15), (a-16), (a-17), (a-18), (a-19), (a-20), (a-21), (a-22), (a-23), (a-24), (a-25), (a-26), (a-27), (a-28), (a-29), (a-31), (a-32), (a-34), (a-35), (a-36), (a-37), (a-38), (a-39), (a-40), (a-41), (a-42) (a-43) or (a-44);

each R⁵ and R⁶ are independently selected from hydrogen; halo; hydroxy; amino; nitro; trihaloC₁₋₆alkyl; trihaloC₁₋₆alkyloxy; C₁₋₆alkyl; C₁₋₆alkyloxy;

C1_6alkyloxyC1_6alkyloxy; C1_6alkylcarbonyl; C1_6alkylsulfonyl; cyanoC1_6alkyl; hydroxyC₁₋₆alkyl; hydroxyC₁₋₆alkyloxy; hydroxyC₁₋₆alkylamino; aminoC₁₋₆alkyloxy; di(C₁₋₆alkyl)aminocarbonyl; di(hydroxyC₁₋₆alkyl)amino; ary[C1_6alkyl)amino; di(C1_6alkyl)aminoC1_6alkyloxy; di(C₁₋₆alkyl)aminoC₁₋₆alkylamino; arylsulfonyl; arylsulfonylamino; aryloxy; arylC2-6alkenediyl; di(C1-6alkyl)amino; di(C_{1.6}alkyl)aminoC_{1.6}alkyl; di(C_{1.6}alkyl)aminoC_{1.6}alkyl(C_{1.6}alkyl)aminoC_{1.6}alkyl; cyano; thiophenyl; thiophenyl substituted with di(C₁₋₆alkyl)aminoC₁₋₆alkyl(C₁₋₆alkyl)aminoC₁₋₆alkyl, di(C₁₋₆alkyl)aminoC₁₋₆alkyl, C₁₋₆alkylpiperazinylC₁₋₆alkyl or di(hydroxyC₁₋₆alkyl)aminoC₁₋₆alkyl; furanyl; imidazolyl; C₁₋₆alkyltriazolyl; tetrazolyl; pyrrolidinyl; piperidinylC₁₋₆alkyloxy; morpholinyl; C_{1.6}alkylmorpholinyl; morpholinylC_{1.6}alkyloxy; morpholinylC₁₋₆alkyl; C₁₋₆alkylpiperazinyl; C₁₋₆alkylpiperazinylC₁₋₆alkyloxy; C_{1-6} alkylpiperazinyl C_{1-6} alkyl; C_{1-6} alkylpiperazinylsulfonyl; aminosulfonylpiperazinylC₁₋₆alkyloxy; aminosulfonylpiperazinyl; aminosulfonylpiperazinylC₁₋₆alkyl; di(C₁₋₆alkyl)aminosulfonylpiperazinyl; di(C₁₋₆alkyl)aminosulfonylpiperazinylC₁₋₆alkyl; hydroxyC₁₋₆alkylpiperazinyl; hydroxyC₁₋₆alkylpiperazinylC₁₋₆alkyl; C₁₋₆alkyloxypiperidinyl; C_{1-6} alkyloxypiperidinyl C_{1-6} alkyl; hydroxy C_{1-6} alkyloxy C_{1-6} alkylpiperazinyl; hydroxyC₁₋₆alkyloxyC₁₋₆alkylpiperazinylC₁₋₆alkyl; (hydroxyC_{1.6}alkyl)(C_{1.6}alkyl)amino; (hydroxyC_{1.6}alkyl)(C_{1.6}alkyl)aminoC_{1.6}alkyl; pyrrolidinylC₁₋₆alkyloxy; pyrazolyl; thiopyrazolyl; pyrazolyl substituted with two substituents selected from C1-6alkyl or trihaloC1-6alkyl; pyridinyl; pyridinyl substituted with C_{1.6}alkyloxy or aryl; pyrimidinyl; quinolinyl; indole; phenyl; phenyl substituted with one, two or three substituents independently selected from halo, amino, C1-6alkyl, C1-6alkyloxy, hydroxyC1-4alkyl, trifluoromethyl, trifluoromethyloxy, hydroxyC₁₋₄alkyloxy, C₁₋₄alkyloxyC₁₋₄alkyloxy, aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)aminoC₁₋₄alkyloxy, di(C₁₋₄alkyl)amino, di(C_{1.4}alkyl)aminoC_{1.4}alkyl, di(C_{1.4}alkyl)aminoC_{1.4}alkyl(C_{1.4}alkyl)aminoC_{1.4}alkyl, piperidinylC₁₋₄alkyloxy, pyrrolidinylC₁₋₄alkyloxy, aminosulfonylpiperazinyl, aminosulfonylpiperazinylC₁₋₄alkyl, di(C₁₋₄alkyl)aminosulfonylpiperazinyl, di(C_{1.4}alkyl)aminosulfonylpiperazinylC_{1.4}alkyl, hydroxyC_{1.4}alkylpiperazinyl, hydroxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, C₁₋₄alkyloxypiperidinyl, C₁₋₄alkyloxypiperidinylC₁₋₄alkyl, hydroxyC₁₋₄alkyloxyC₁₋₄alkylpiperazinyl, hydroxyC₁₋₄alkyloxyC₁₋₄alkylpiperazinylC₁₋₄alkyl, (hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)amino, (hydroxyC₁₋₄alkyl)(C₁₋₄alkyl)aminoC₁₋₄alkyl, pyrrolidinylC₁₋₄alkyloxy, morpholinylC₁₋₄alkyloxy, morpholinylC₁₋₄alkyl,

 $\label{eq:continuous_continuous_continuous} C_{1\text{-4}alkylpiperazinyl} C_$

292. The compound of claim 287 that is selected from one of

wherein the terminal hydroxamic acid moiety (-C(O)-NH-OH) is replaced with

wherein Φ , R^1 , R^2 , R^3 , and R^4 are as defined in accordance with claim 1.

- 293. The compound of claim 287 wherein R¹, R², R³, and R⁴ are all H.
- 294. A compound according to claim 287 for use in inhibting histone deacetylase.
- 295. A compound according to calim 287 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.
- 296. The compound of claim 295, wherein said treatment is effected by inhibiting histone deacetylase.
- 297. The compound of calim 295, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 298. The compound of claim 295, wherein said cell proliferative disease is cancer.
- 299. The compound of claim 398, wherein said cancer is a solid tumor cancer.
- 300. The compound of claim 298, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 301. A pharmaceutical composition comprising a compound according to claim 287 and a pharmaceutically acceptable carrier.
- 302. The pharmaceutical composition of claim 301 further comprising a nucleic acid level inhibitor of histone deacetylase.
- 303. The pharmaceutical composition of claim 302, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 304. The pharmaceutical composition of claim 303, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4,

SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.

- 305. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 287.
- 306. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 301.
- 307. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 302.
- 308. The method of claim 306, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 309. The method of claim 306, wherein said cell proliferative disease is cancer.
- 310. The method of claim 309, wherein said cancer is a solid tumor cancer.
- 311. The method of claim 310, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 312. The method of claim 307, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 313. The method of claim 30, wherein said cell proliferative disease is cancer.
- 314. The method of claim 313, wherein said cancer is a solid tumor cancer.
- 315. The method of claim 314, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 316. A compound of the formula:

or a pharmaceutically acceptable salt thereof, wherein

 Φ is -NH₂ or -OH;

R1 is H or as defined in claim 1:

R², R³, and R⁴ are as defined in claim 1:

Ring A is a heterocyclyl, wherein if said heterocyclyl contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from G;

R¹¹ is a substituent on carbon and is selected from halo, nitro, cyano, hydroxy, oxo, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₁₋₆alkoxy, C₁₋₆alkanoyl, C₁₋₆alkanoyloxy, N-(C₁₋₆alkyl)amino, N,N-(C₁₋₆alkyl)₂amino, C₁₋₆alkanoylamino, N-(C₁₋₆alkyl)carbamoyl, N,N-(C₁₋₆alkyl)₂carbamoyl, C₁₋₆alkylS(O)_a wherein a is 0 to 2, C₁₋₆alkoxycarbonyl, N-(C₁₋₆alkyl)₂sulphamoyl, aryl, aryloxy, arylC₁₋₆alkyl, heterocyclic group, (heterocyclic group)C₁₋₆alkyl or a group (D-E-); wherein R¹, including group (D-E-), may be optionally substituted on carbon by one or more V; and wherein, if said heterocyclic group contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from J;

V is halo, nitro, cyano, hydroxy, oxo, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{1-6} alkoxy, C_{1-6} alkanoyl, C_{1-6} alkanoyloxy, $N-(C_{1-6}$ alkyl)amino, $NN-(C_{1-6}$ alkyl)2amino, C_{1-6} alkanoylamino, $N-(C_{1-6}$ alkyl)2arbamoyl, $NN-(C_{1-6}$ alkyl)2carbamoyl, C_{1-6} alkyl)2carbamoyl, $NN-(C_{1-6}$ alkyl)2sulphamoyl wherein a is 0 to 2, C_{1-6} alkoxycarbonyl, $N-(C_{1-6}$ alkyl)3sulphamoyl, $NN-(C_{1-6}$ alkyl)2sulphamoyl

or a group (D'-E'-); wherein V, including group (D'-E'-), may be optionally substituted on carbon by one or more W;

W and Z are independently selected from halo, nitro, cyano, hydroxy, oxo, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₁₋₆alkoxy, C₁₋₆alkanoyl, C₁₋₆alkanoyloxy, N-(C₁₋₆alkyl)amino, N,N-(C₁₋₆alkyl)₂amino, C₁₋₆alkanoylamino, N-(C₁₋₆alkyl)carbamoyl, N,N-(C₁₋₆alkyl)₂carbamoyl, C₁₋₆alkylS(O)_a wherein a is 0 to 2, C₁₋₆alkoxycarbonyl, N-(C₁₋₆alkyl)₃sulphamoyl or N,N-(C₁₋₆alkyl)₂sulphamoyl;

G, J and K are independently selected from C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkanoyl, C₁₋₈alkylsulphonyl, C₁₋₈alkoxycarbonyl, carbamoyl, N-(C₁₋₈alkyl)carbamoyl, N,N-(C₁₋₈alkyl)carbamoyl, benzyloxycarbonyl, benzoyl and phenylsulphonyl, aryl, arylC₁₋₆alkyl or (heterocyclic group)C₁₋₆alkyl; wherein G, J and K may be optionally substituted on carbon by one or more Q; and wherein if said heterocyclic group contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from hydrogen or C₁₋₆alkyl;

Q is halo, nitro, cyano, hydroxy, oxo, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₁₋₆alkoxy, C₁₋₆alkanoyl, C₁₋₆alkanoyloxy, N-(C₁₋₆alkyl)amino, N,N-(C₁₋₆alkyl)₂amino, C₁₋₆alkanoylamino, N-(C₁₋₆alkyl)carbamoyl, N,N-(C₁₋₆alkyl)₂carbamoyl, C₁₋₆alkylS(O)_a wherein a is 0 to 2, C₁₋₆alkoxycarbonyl, C₁₋₆alkoxycarbonylamino, N-(C₁₋₆alkyl)sulphamoyl, N,N-(C₁₋₆alkyl)₂sulphamoyl, aryl, aryloxy, arylC₁₋₆alkyl, arylC₁₋₆alkoxy, heterocyclic group, (heterocyclic group)C₁₋₆alkyl, (heterocyclic group)C₁₋₆alkoxy, or a group (D"-E"-); wherein Q, including group (D"-E"-), may be optionally substituted on carbon by one or more Z;

- D, D' and D'' are independently selected from C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl,
 C₃₋₈cycloalkyl, C₃₋₈cycloalkylC₁₋₆alkyl, aryl, arylC₁₋₆alkyl, heterocyclic group, (heterocyclic group)C₁₋₆alkyl; wherein D, D' and D'' may be optionally substituted on carbon by one or more F'; and wherein if said heterocyclic group contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from K;
- E, E' and E'' are independently selected from -N(R^a)-, -O-, -C(O)O-, -OC(O)-,

 -C(O)-, -N(R^a)C(O)-, -N(R^a)C(O)N(R^b)-, -N(R^a)C(O)O-, -OC(O)N(R^a)-, -C(O)N(R^a)-,

 -S(O)_r-, -SO₂N(R^a)-, -N(R^a)SO₂-; wherein R^a and R^bare independently selected from hydrogen or C₁₋₆alkyl optionally substituted by one or more F and r is 0-2;

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F and F' are independently selected from halo, nitro, cyano, hydroxy, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, C_{1-6} alkoxy, C_{1-6} alkanoyl, C_{1-6} alkanoyloxy, N-(C_{1-6} alkyl)amino,

 $N,N-(C_{1-6}alkyl)_2$ amino, $C_{1-6}alkanoylamino$, $N-(C_{1-6}alkyl)$ carbamoyl,

 $N,N-(C_{1-6}alkyl)_2$ carbamoyl, $C_{1-6}alkylS(O)_a$ wherein a is 0 to 2, $C_{1-6}alkoxycarbonyl$, $N-(C_{1-6}alkyl)_2$ sulphamoyl;

m is 0, 1, 2, 3 or 4; wherein the values of R¹ may be the same or different; Ring B is a ring selected from

wherein,

X1 and X2 are selected from CH or N, and

 Y^1 , Y^2 , Y^3 and Y^4 are selected from CH or N provided that at least one of Y^1 , Y^2 , Y^3 and Y^4 is N;

R¹² is halo:

n is 0, 1, or 2, wherein the values of R¹² are the same or different.

- 317. The compound of claim 316 wherein
 - Ring A is a pyridyl, quinolyl, indolyl, pyrimidinyl, morpholinyl, piperidinyl, piperazinyl, pyridazinyl, pyrazinyl, thiazolyl, thienyl, thienopyrimidinyl, thienopyridinyl, purinyl, 1',2',3',6'-tetrahydropyridinyl, triazinyl, oxazolyl, pyrazolyl, or furanyl; wherein if Ring A contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from G.
 - Ring B is thienyl, thiadiazolyl, thiazolyl, pyrimidyl, pyrazinyl, pyridazinyl or pyridyl.
 - or
 - Ring B is thienyl or pyridyl wherein both the thienyl and the pyridyl are attached to Ring A in the 2-position of the thienyl or pyridyl ring and to the amide group of formula (I) in the 5-position of the thienyl or pyridyl ring.

• is halo, amino, C₁₋₆alkyl, C₁₋₆alkoxy, C₁₋₃alkanoyloxy, N-(C₁₋₃alkyl)amino, N,N-(C₁₋₃alkyl)₂amino, C₁₋₃alkanoylamino, N-(C₁₋₃alkyl)₂carbamoyl, N,N-(C₁₋₃alkyl)₂carbamoyl.

- or
- R¹¹ is a substituent on carbon and is selected from halo, nitro, cyano, hydroxy, oxo, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₁₋₆alkoxy, C₁₋₆alkanoyl, C₁₋₆alkanoyloxy, N-(C₁₋₆alkyl)amino, N,N-(C₁₋₆alkyl)₂amino, C₁₋₆alkanoylamino, N-(C₁₋₆alkyl)carbamoyl, N,N-(C₁₋₆alkyl)₂carbamoyl, C₁₋₆alkylS(O)_n wherein a is 0 to 2, C₁₋₆alkoxycarbonyl, N-(C₁₋₆alkyl)sulphamoyl, N,N-(C₁₋₆alkyl)₂sulphamoyl, aryl, aryloxy, arylC₁₋₆alkyl, heterocyclic group, (heterocyclic group)C₁₋₆alkyl or a group (D-E-); wherein R¹, including group (D-E-), may be optionally substituted on carbon by one or more V; and wherein, if said heterocyclic group contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from J;

V is halo, nitro, cyano, hydroxy, oxo, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₁₋₆alkoxy, C₁₋₆alkanoyl, C₁₋₆alkanoyloxy, N-(C₁₋₆alkyl)amino, N,N-(C₁₋₆alkyl)₂amino,

C₁₋₆alkanoylamino, N-(C₁₋₆alkyl)carbamoyl, N,N-(C₁₋₆alkyl)₂carbamoyl, C₁₋₆alkylS(O)_a wherein a is 0 to 2, C₁₋₆alkoxycarbonyl, N-(C₁₋₆alkyl)sulphamoyl, N,N-(C₁₋₆alkyl)₂sulphamoyl or a group (D'-E'-); wherein V, including group (D'-E'-), may be optionally substituted on carbon by one or more W;

W and Z are independently selected from halo, nitro, cyano, hydroxy, oxo, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkyr.yl, C₁₋₆alkoxy, C₁₋₆alkanoyl, C₁₋₆alkanoyloxy, N-(C₁₋₆alkyl)amino, N,N-(C₁₋₆alkyl)₂amino, C₁₋₆alkanoylamino, N-(C₁₋₆alkyl)carbamoyl, N,N-(C₁₋₆alkyl)₂carbamoyl, C₁₋₆alkylS(O)_a wherein a is 0 to 2, C₁₋₆alkoxycarbonyl, N-(C₁₋₆alkyl)sulphamoyl or N,N-(C₁₋₆alkyl)₂sulphamoyl;

G, J and K are independently selected from C_{1-8} alkyl, C_{2-8} alkenyl, C_{2-8} alkynyl, C_{1-8} alkylsulphonyl, C_{1-8} alkoxycarbonyl, carbamoyl, N-(C_{1-8} alkyl)carbamoyl, benzyloxycarbonyl, benzoyl and phenylsulphonyl, aryl, aryl C_{1-6} alkyl or (heterocyclic group) C_{1-6} alkyl; wherein G, J and K may be optionally substituted on carbon by one or more Q; and wherein if said heterocyclic group contains an - NH- moiety that nitrogen may be optionally substituted by a group selected from hydrogen or C_{1-6} alkyl;

Q is halo, nitro, cyano, hydroxy, oxo, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₁₋₆alkoxy, C₁₋₆alkanoyl, C₁₋₆alkanoyloxy, N-(C₁₋₆alkyl)amino, N,N-(C₁₋₆alkyl)₂amino, C₁₋₆alkanoylamino, N-(C₁₋₆alkyl)carbamoyl, N,N-(C₁₋₆alkyl)₂carbamoyl, C₁₋₆alkylS(O)₂ wherein a is 0 to 2, C₁₋₆alkoxycarbonyl, C₁₋₆alkoxycarbonylamino, N-(C₁₋₆alkyl)sulphamoyl, N,N-(C₁₋₆alkyl)₂sulphamoyl, aryl, aryloxy, arylC₁₋₆alkyl, arylC₁₋₆alkoxy, heterocyclic group, (heterocyclic group)C₁₋₆alkyl, (heterocyclic group)C₁₋₆alkoxy, or a group (D"-E"-); wherein Q, including group (D"-E"-), may be optionally substituted on carbon by one or more Z;

D, D' and D'' are independently selected from C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₃₋₈cycloalkyl, C₃₋₈cycloalkylC₁₋₆alkyl, aryl, arylC₁₋₆alkyl, heterocyclic group, (heterocyclic group)C₁₋₆alkyl; wherein D, D' and D'' may be optionally substituted on carbon by one or more F'; and wherein if said heterocyclic group contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from K:

E, E' and E'' are independently selected from -N(R^a)-, -O-, -C(O)O-, -OC(O)-, -C(O)-, -N(R^a)C(O)-, -N(R^a)C(O)N(R^b)-, -N(R^a)C(O)O-, -OC(O)N(R^a)-, -C(O)N(R^a)-, -S(O),-,

 $-SO_2N(R^a)$ -, $-N(R^a)SO_2$ -; wherein R^a and R^b are independently selected from hydrogen or C_{1-6} alkyl optionally substituted by one or more F and r is 0-2; and

F and F' are independently selected from halo, nitro, cyano, hydroxy, trifluoromethyl, trifluoromethoxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C₁₋₆alkyl, C₂₋₆alkenyl, C₂₋₆alkynyl, C₁₋₆alkoxy, C₁₋₆alkanoyl, C₁₋₆alkanoyloxy, N-(C₁₋₆alkyl)amino, N,N-(C₁₋₆alkyl)₂amino, C₁₋₆alkanoylamino, N-(C₁₋₆alkyl)carbamoyl, N,N-(C₁₋₆alkyl)₂carbamoyl, C₁₋₆alkylS(O)_a wherein a is 0 to 2, C₁₋₆alkoxycarbonyl, N-(C₁₋₆alkyl)₂sulphamoyl and N,N-(C₁₋₆alkyl)₂sulphamoyl.

m is 0, 1, 2, 3 or 4; wherein the values of R¹¹ are the same or different.

- R¹² is halo.
- n is 0, 1, or 2; wherein the values of R¹² are the same or different;
- 318. The compound of claim 317 wherein

• Ring A is pyridin-4-yl, pyridin-3-yl, pyridin-2-yl, quinolin-8-yl, pyrimidin-6-yl, pyrimidin-5-yl, pyrimidin-4-yl, morpholin-4-yl, piperidin-4-yl, piperidin-3-yl, piperdin-2-yl, piperazin-4-yl, pyridazin-5-yl, pyrazin-6-yl, thiazol-2-yl, thiazol-2-yl,

Ring B is thienyl, thiazolyl, pyrimidyl, pyrazinyl, pyridazinyl or pyridyl.

• R¹¹ is halo, amino, C₁₋₆alkyl or C₁₋₆alkoxy.

319. The compound of claim 317 wherein

Ring A is pyridin-4-yl, pyridin-3-yl, pyridin-2-yl, morpholin-4-yl, piperidin-4-yl, piperidin-3-yl, piperidin-2-yl, piperazin-4-yl, thiazol-2-yl, thien-2-yl, furan-3-yl, pyrrolidin-1-yl, piperidin-1-yl, triazol-1-yl or 1',2',3',6'-tetrahydropyridin-4-yl wherein if Ring A contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from G.

- Ring B is thienyl or pyridyl.
- R¹¹ is halo, amino, methyl or methoxy.
- 320. The compound of claim 317 wherein

Ring A is a pyridyl, pyrimidyl, morpholinyl, piperidinyl, piperazinyl, pyridazinyl, thienyl, pyrazinyl, thiazolyl, 1,2,4-triazolyl or furanyl.

321. The compound of claim 317 wherein

Ring A is pyridin-4-yl, pyridin-3-yl, pyridin-2-yl or 1,2,4-triazolyl.

322. The compound of claim 317 wherein

substituent on carbon and is selected from cyano, hydroxy, C₁₋₆alkyl or a group (D-E-); who wherein R¹¹ including group E-), may be optionally substituted on carbon by one or more V;

V is cyano, hydroxy or a group (D'-E'-); wherein V, including group (D'-E'-), may be optionally substituted on carbon by one or more W;

W and Z are independently selected from cyano, C_{1.6}alkyl or C_{1.6}alkoxy;

G and K are independently selected from C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, arylC₁₋₆alkyl or (heterocyclic group)C₁₋₆alkyl; wherein G and K may be optionally substituted on carbon by one or more Q;

Q is cyano, hydroxy, oxo, C₁₋₆alkyl, C₂₋₆alkenyl, C₁₋₆alkoxy, C₁₋₆alkanoyl, C₁₋₆alkanoyloxy, N-(C₁₋₆alkyl)carbamoyl, N,N-(C₁₋₆alkyl)₂carbamoyl, C₁₋₆alkoxycarbonyl, C₁₋₆alkoxycarbonylamino, aryl, aryloxy or a group (D"-B'-); wherein Q, including group (D"-E"-), may be optionally substituted on carbon by one or more Z;

D, D' and D'' are independently selected from aryl, arylC₁₋₆alkyl or heterocyclic group; wherein D, D' and D'' may be optionally substituted on carbon by one or more F'; and wherein if said heterocyclic group contains an -NH- moiety that nitrogen may be optionally substituted by a group selected from K;

E, E' and B'' are independently selected from -O-, -C(O)O-, -OC(O)-, -C(O)-, -N(R^a)C(O)-, -C(O)N(R^a)-, -S(O)_r; wherein R^a is selected from hydrogen or C_{1-6} alkyl optionally substituted by one or more F and r is 0-2; and

F and F' are independently selected from nitro, hydroxy, C_{1-6} alkyl, C_{1-6} alkoxy, C_{1-6} alkanoyl, N- $(C_{1-6}$ alkyl)amino, N, N- $(C_{1-6}$ alkyl)₂amino, C_{1-6} alkanoylamino or C_{1-6} alkoxycarbonyl.

- 323. The compound of claim 317 wherein R¹² is fluoro.
- 324. The compound of claim 317 wherein R¹² is chloro.
- 325. The compound of claim 316 wherein each of R^2 , R^3 , and R^4 corresponds to R^{12} , R^{13} , and R^{14} , respectively, in claim 316 wherein:

Ring A is a pyridyl, indolyl, pyrimidyl, morpholinyl, piperidinyl, piperazinyl, pyridazinyl, thienyl, pyrazinyl, thiazolyl, oxazolyl, 1,2,4-triazolyl, isoxazolyl, isothiazolyl, pyrazolyl, orfuranyl;

Ring B is thienyl, thiadiazolyl, thiazolyl, pyiitnidyl, pyrazinyl, pyridazinyl or pyridyl;

 R^{11} is halo, amino, C_{1-1} alkyl, C_{1-1} alko xy, C_{1-1} alkanoyloxy, $N-(C_{1-1}$ alky $I)_2$ amino,

 N_{i} N-(C_{1-3} lky I)₂amino, C_{1-3} alkanoylamino, N-(C_{1-3} alkyl)carbamoyl, N_{i} N-(C_{1-4} lky)₂carbamoyl;

m is 0, 1, 2, wherein the values of R11 are the same or different.

n is 0, 1, 2, wherein the values of R12 are the same or different;

R12 is F or Cl.

326. The compound of claim 316 wherein each of \mathbb{R}^2 , \mathbb{R}^3 , and \mathbb{R}^4 corresponds to \mathbb{R}^{12} , \mathbb{R}^{13} , and \mathbb{R}^{14} , respectively, in claim 316 wherein:

Ring A is pyridin-4-yI, pyridin-3-yl, pyridin-2-yl or 1,2,4-triazolyl;

Ring B is thienyl or pyridyl;

R¹¹ is halo, amino, methyl or methoxy;

m is 0, 1, 2, wherein the values of R11 are the same or different,

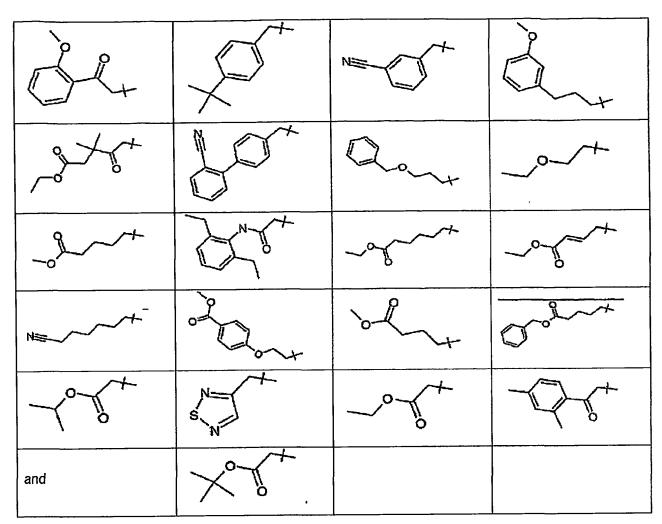
n is 0 or 1;

R12 is F.

327. The compound of claim 316 that is

wherein R¹¹ is selected from one of:

			**
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0	+>-	Dort	Only
of M Done	T	2-4	Xolunt
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Don	201	°D on	9-Q-4



328. The compound of claim 316 wherein \mathbb{R}^2 , \mathbb{R}^3 , and \mathbb{R}^4 are all H.

329. The compound of claim 316 that is selected from one of the compounds of WO 03/024448 wherein the terminal moieties -C(O)-NH-Ay¹, -C(O)-NH-Ay², -C(O)-NH-Ar^a-NH₂, and

are replaced with the moiety:

$$\begin{array}{c|c}
R^1 \\
R^2 \\
R^3 \\
R^4
\end{array}$$

wherein Φ, R1, R2, R3, and R4 are as defined in accordance with claim 1.

- 330. A compound according to claim 316 for use in inhibting histone deacetylase.
- 331. A compound according to calim 316 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.
- 332. The compound of claim 331, wherein said treatment is effected by inhibiting histone deacetylase.
- 334. The compound of calim 331, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 335. The compound of claim 331, wherein said cell proliferative disease is cancer.
- 336. The compound of claim 335, wherein said cancer is a solid tumor cancer.
- 337. The compound of claim 335, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 338. A pharmaceutical composition comprising a compound according to claim 316 and a pharmaceutically acceptable carrier.
- 339. The pharmaceutical composition of claim 338 further comprising a nucleic acid level inhibitor of histone deacetylase.
- 340. The pharmaceutical composition of claim 339, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 341. The pharmaceutical composition of claim 340, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1 SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.

342. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 316.

- 343. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 338.
- 344. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 339.
- 345. The method of claim 343, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 346. The method of claim 343, wherein said cell proliferative disease is cancer.
- 347. The method of claim 346, wherein said cancer is a solid tumor cancer.
- 348. The method of claim 347, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 349. The method of claim 344, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 350. The method of claim 344, wherein said cell proliferative disease is cancer.
- 351. The method of claim 350, wherein said cancer is a solid tumor cancer.
- 352. The method of claim 351, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 353. A compound of the formula:

Ar-A-D-E-G-NH-
$$\mathbb{R}^1$$
 \mathbb{R}^2
 \mathbb{R}^2
 \mathbb{R}^3

or a pharmaceutically acceptable salt thereof, wherein

 Φ is -NH $_2$ or -OH;

R1 is H or as defined in claim 1;

R2, R3, and R4 are as defined in claim 1; and

Ar, A, D, E, and G are as defined in JP 2003137866.

354. The compound of claim 353 wherein R1, R2, R3, and R4 are all H.

355. The compound of claim 353 that is selected from one of the compounds of JP 2003137866 wherein the terminal moiety:

wherein Φ, R1, R2, R3, and R4 are as defined in accordance with claim 1.

356. A compound according to claim 353 for use in inhibting histone deacetylase.

- 357. A compound according to calim 353 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.
- 358. The compound of claim 357, wherein said treatment is effected by inhibiting histone deacetylase.
- 359. The compound of calim 357, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 360. The compound of claim 357, wherein said cell proliferative disease is cancer.
- 361. The compound of claim 360, wherein said cancer is a solid tumor cancer.
- 362. The compound of claim 360, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 363. A pharmaceutical composition comprising a compound according to claim 353 and a pharmaceutically acceptable carrier.
- 364. The pharmaceutical composition of claim 363 further comprising a nucleic acid level inhibitor of histone deacetylase.

365. The pharmaceutical composition of claim 364, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.

- 366. The pharmaceutical composition of claim 365, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1 SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.
- 367. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 353.
- 368. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 363.
- 369.A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 364.
- 370. The method of claim 368, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 371. The method of claim 368, wherein said cell proliferative disease is cancer.
- 372. The method of claim 371, wherein said cancer is a solid tumor cancer.
- 373. The method of claim 372, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 374. The method of claim 369, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 375. The method of claim 369, wherein said cell proliferative disease is cancer.
- 376. The method of claim 375, wherein said cancer is a solid tumor cancer.

377. The method of claim 376, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.

378. A compound of the formula:

or a pharmaceutically acceptable salt thereof, wherein

O is -NH 2 or -OH;

R1 is H or as defined in claim 1;

R², R³, and R⁴ are as defined in claim 1;

X, Y, and A are as defined in JP 11-269146 (1999); and

R¹¹ is the same as R¹ of JP 11-269146 (1999).

379. The compound of claim 378 wherein R^1 , R^2 , R^3 , and R^4 are all H.

380. The compound of claim 378 that is selected from one of the compounds 1-50 of Tables 2-4 of JP 11-269146 (1999) wherein the terminal moiety:

wherein Φ , R^1 , R^2 , R^3 , and R^4 are as defined in accordance with claim 1.

381. A compound according to claim 378 for use in inhibting histone deacetylase.

- 382. A compound according to calim 378 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.
- 383. The compound of claim 382, wherein said treatment is effected by inhibiting histone deacetylase.

384. The compound of calim 382, wherein said cell proliferative disease is a neoplastic cell proliferative disease.

- 385. The compound of claim 382, wherein said cell proliferative disease is cancer.
- 386. The compound of claim 385, wherein said cancer is a solid tumor cancer.
- 387. The compound of claim 385, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 388. A pharmaceutical composition comprising a compound according to claim 378 and a pharmaceutically acceptable carrier.
- 389. The pharmaceutical composition of claim 388 further comprising a nucleic acid level inhibitor of histone deacetylase.
- 390. The pharmaceutical composition of claim 389, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 391. The pharmaceutical composition of claim 390, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.
- 392. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 378.
- 393. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 388.
- 394. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 389.
- 395. The method of claim 393, wherein said cell proliferative disease is a neoplastic cell proliferative disease.

- 396. The method of claim 393, wherein said cell proliferative disease is cancer.
- 397. The method of claim 396, wherein said cancer is a solid tumor cancer.
- 398. The method of claim 397, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 399. The method of claim 394, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 400. The method of claim 394, wherein said cell proliferative disease is cancer.
- 401. The method of claim 400, wherein said cancer is a solid tumor cancer.
- 402. The method of claim 401, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 403. A compound of the formula:

$$A-X-Q-(CH_2)_n$$
 R^{11}
 O
 NH
 R^2
 R^3

or a pharmaceutically acceptable salt thereof, wherein

O is -NH 2 or -OH;

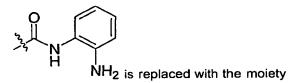
R1 is H or as defined in claim 1;

R2, R3, and R4 are as defined in claim 1;

n, X, Q, and A are as defined in JP 11-302173 (1999); and

R¹¹ is the same as R¹ of JP 11-302173 (1999).

- 404. The compound of claim 403 wherein R¹, R², R³, and R⁴ are all H.
- 405. The compound of claim 403 that is selected from one of the compounds 1-67 of JP 11-302173 (1999) wherein the terminal moiety:



$$\begin{array}{c|c}
 & R^1 \\
 & R^2 \\
 & R^3 \\
 & R^4 \\
 & R^4
\end{array}$$

wherein Φ , R^1 , R^2 , R^3 , and R^4 are as defined in accordance with claim 1.

- 406. A compound according to claim 403 for use in inhibting histone deacetylase.
- 407. A compound according to calim 403 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.
- 408. The compound of claim 407, wherein said treatment is effected by inhibiting histone deacetylase.
- 409. The compound of calim 407, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 410. The compound of claim 407, wherein said cell proliferative disease is cancer.
- 411. The compound of claim 410, wherein said cancer is a solid tumor cancer.
- 412. The compound of claim 410, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 413. A pharmaceutical composition comprising a compound according to claim 403 and a pharmaceutically acceptable carrier.
- 414. The pharmaceutical composition of claim 413 further comprising a nucleic acid level inhibitor of histone deacetylase.
- 415. The pharmaceutical composition of claim 414, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 416. The pharmaceutical composition of claim 415, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.

417. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 403.

- 418. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 413.
- 419. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 414.
- 420. The method of claim 418, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 421. The method of claim 418, wherein said cell proliferative disease is cancer.
- 422. The method of claim 421, wherein said cancer is a solid tumor cancer.
- 423. The method of claim 422, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 424. The method of claim 419, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 425. The method of claim 419, wherein said cell proliferative disease is cancer.
- 426. The method of claim 425, wherein said cancer is a solid tumor cancer.
- 427. The method of claim 426, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 428. A compound of the formula:

$$A-X-Q-(CH_2)_n$$
 R^{11}
 O
 NH
 R^2
 R^3

or a pharmaceutically acceptable salt thereof, wherein

 Φ is -NH $_2$ or -OH;

R¹ is H or as defined in claim 1;

 R^2 , R^3 , and R^4 are as defined in claim 1;

n, Q, and A are as defined in JP 2001 131 130; and

R¹¹ is the same as R¹ of JP 2001131130.

- 429. The compound of claim 428 wherein R¹, R², R³, and R⁴ are all H.
- 430. The compound of claim 428 that is selected from one of the compounds of JP 2001131130 wherein the terminal moieties

NH₂ and NH₂ are replaced with the moiety
$$\mathbb{R}^1$$
 \mathbb{R}^2 \mathbb{R}^3 \mathbb{R}^4

wherein Φ, R¹, R², R³, and R⁴ are as defined in accordance with claim 1.

- 431. A compound according to claim 428 for use in inhibting histone deacetylase.
- 432. A compound according to calim 428 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.
- 433. The compound of claim 432, wherein said treatment is effected by inhibiting histone deacetylase.
- 434. The compound of calim 432, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 435. The compound of claim 432, wherein said cell proliferative disease is cancer.
- 436. The compound of claim 435, wherein said cancer is a solid tumor cancer.
- 437. The compound of claim 435, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 438. A pharmaceutical composition comprising a compound according to claim 428 and a pharmaceutically acceptable carrier.

439. The pharmaceutical composition of claim 438 further comprising a nucleic acid level inhibitor of histone deacetylase.

- 440. The pharmaceutical composition of claim 439, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 441. The pharmaceutical composition of claim 440, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:I, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.
- 442. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 428.
- 443. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 438.
- 444. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 439.
- 445. The method of claim 443, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 446. The method of claim 443, wherein said cell proliferative disease is cancer.
- 447. The method of claim 446, wherein said cancer is a solid tumor cancer.
- 448. The method of claim 447, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 449. The method of claim 444, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 450. The method of claim 444, wherein said cell proliferative disease is cancer.
- 451. The method of claim 450, wherein said cancer is a solid tumor cancer.

452. The method of claim 451, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.

453. A compound of formula:

$$A-X-Q-(CH_2)_n$$
 R^{11}
 O
 NH
 R^2
 R^3

or a pharmaceutically acceptable salt thereof, wherein

 Φ is -NH $_2$ or -OH;

R1 is H or as defined in claim 1;

R², R³, and R⁴ are as defined in claim 1;

n, X, Q, and A are as defined in JP 10152462, JP 2002332267, and JP 11-302173; and R^{11} is the same as R^1 of JP 10152462, JP 2002332267, and JP 11-302173.

454. The compound of claim 453 wherein R¹, R², R³, and R⁴ are all H.

455. The compound of claim 453 that is selected from one of the compounds of JP 10152462, JP 2002332267, and JP 11-302173 wherein the terminal moiety

is replaced with the moiety:

$$\begin{array}{c|c} & & & R^1 \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & & \\$$

wherein Φ , R^1 , R^2 , R^3 , and R^4 are as defined in accordance with claim 1.

456. A compound according to claim 453 for use in inhibting histone deacetylase.

- 457. A compound according to calim 453 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.
- 458. The compound of claim 457 wherein said treatment is effected by inhibiting histone deacetylase.

459. The compound of calim 457, wherein said cell proliferative disease is a neoplastic cell proliferative disease.

- 460. The compound of claim 457, wherein said cell proliferative disease is cancer.
- 461. The compound of claim 460, wherein said cancer is a solid tumor cancer.
- 462. The compound of claim 460, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 463. A pharmaceutical composition comprising a compound according to claim 453 and a pharmaceutically acceptable carrier.
- 464. The pharmaceutical composition of claim 463 further comprising a nucleic acid level inhibitor of histone deacetylase.
- 465. The pharmaceutical composition of claim 464, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 466. The pharmaceutical composition of claim 465, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4, SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.
- 467. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 453.
- 468. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 463.
- 469. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 464.
- 470. The method of claim 468, wherein said cell proliferative disease is a neoplastic cell proliferative disease.

- 471. The method of claim 468, wherein said cell proliferative disease is cancer.
- 472. The method of claim 471, wherein said cancer is a solid tumor cancer.
- 473. The method of claim 472, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 474. The method of claim 469, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 475. The method of claim 469, wherein said cell proliferative disease is cancer.
- 476. The method of claim 475, wherein said cancer is a solid tumor cancer.
- 477. The method of claim 476, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 478. A compounds of the formula:

$$A-X-Q-(CH_2)_n$$
 R^{11}
 O
 NH
 R^2
 R^3

or a pharmaceutically acceptable salt thereof, wherein

 Φ is -NH₂ or -OH;

R1 is H or as defined in claim 1;

R², R³, and R⁴ are as defined in claim 1;

n, X, Q, and A are as defined in US 6,174,905; and

R¹¹ is the same as R¹ of US 6,174,905.

- 479. The compound of claim 478 wherein R1, R2, R3, and R4 are all H.
- 480. The compound of claim 478 that is selected from one of the compounds of US
- 6,174,905 wherein the terminal moiety:

of the compounds of Table 1 of US 6,174,905 and the terminal moiety:

of the compounds of Tables 2-4 of US 6,174,905 are replaced with the moiety:

$$\begin{array}{c} \begin{array}{c} R^1 \\ R^2 \\ \end{array}$$

wherein Φ , R^1 , R^2 , R^3 , and R^4 are as defined in accordance with paragraph claim 1.

- 481. A compound according to claim 478 for use in inhibting histone deacetylase.
- 482. A compound according to calim 478 for use in treatment of a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease.
- 483. The compound of claim 482, wherein said treatment is effected by inhibiting histone deacetylase.
- 484. The compound of calim 482, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 485. The compound of claim 482, wherein said cell proliferative disease is cancer.
- 486. The compound of claim 485, wherein said cancer is a solid tumor cancer.
- 487. The compound of claim 485, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 488. A pharmaceutical composition comprising a compound according to claim 478 and a pharmaceutically acceptable carrier.
- 489. The pharmaceutical composition of claim 488 further comprising a nucleic acid level inhibitor of histone deacetylase.
- 490. The pharmaceutical composition of claim 489, wherein siad nucleic acid level inhibitor is an antisense oligonucleotide complementary to a nucleic acid that encodes for a histone deacetylase.
- 491. The pharmaceutical composition of claim 490, wherein said antisense oligonucleotide is selected from the group consisting if SEQ ID No:1, SEQ ID No:2, SEQ ID No:3, SEQ ID No:4,

SEQ ID No:5, SEQ ID No:6, SEQ ID No:7, SEQ ID No:8, SEQ ID No:9, SEQ ID No:10, SEQ ID No:11, SEQ ID No:12, SEQ ID No:13, SEQ ID No:14, SEQ ID No:15, SEQ ID No:16, and SEQ ID No:17.

- 492. A method of inhibiting histone deacetylase, the method comprising contacting said histone deacetylase with an inhibiting efective amount of a compound according to claim 478.
- 493. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 488.
- 494. A method of treating an individual having a disease selected from the group consisting of a cell proliferative disease, a protozoal disease and a fungal disease, said method comprising administering to said individual a treatment effective amount of the pharmaceutical composition of claim 489.
- 495. The method of claim 493, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 496. The method of claim 493, wherein said cell proliferative disease is cancer.
- 497. The method of claim 496, wherein said cancer is a solid tumor cancer.
- 498. The method of claim 497, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- 499. The method of claim 494, wherein said cell proliferative disease is a neoplastic cell proliferative disease.
- 500. The method of claim 494, wherein said cell proliferative disease is cancer.
- 501. The method of claim 500, wherein said cancer is a solid tumor cancer.
- 502. The method of claim 501, wherein said cancer is selected from the group consisting of a lymphoma, lung cancer, colon cancer, prostrate cancer, stomach cancer, breast cancer and leukemia.
- A compound selected from the compounds of Table 1 and Table I a and pharmaceutically acceptable salts thereof.